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***Introducing and Evaluating Happiness as a New Patient  
Reported Outcome in Dermatology***

vorgelegt von:

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aus:

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I hereby declare, that the submitted thesis entitled:

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is my own work. I have only used the sources indicated and have not made unauthorised use of services of a third party. Where the work of others has been quoted or reproduced, the source is always given.

I further declare that the submitted thesis or parts thereof have not been presented as part of an examination degree to any other university.

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## Confirmation of congruency



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## List of abbreviations

DLQI: Dermatology Quality of Life Index

EMA: European Medicines Agency

FDA: (American) Food and Drug Administration

ICD: International Classification of Diseases

HIV: human immunodeficiency virus

m<sup>a</sup>: adjusted mean

NA: negative affect

PA: positive affect

PANAS: Positive and Negative Affect Schedule

PROM: patient reported outcome measure

QoL: Quality of Life

SPANE: Scale of Positive and Negative Experience

SWL: satisfaction with life

SWLS: Satisfaction With Life Scale

WHO: World Health Organization

## List of publications

**Schuster B**, Ziefreund S, Schielein MC, Tizek L, Biedermann T, Peifer C, Zink A. Adding happiness to complement the Dermatology Quality of Life Index in psoriasis and atopic dermatitis healthcare. *Eur J Dermatol*. Accepted.

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# **1. Your contribution to the publications**

## **1.1 Contribution to paper I**

I, Barbara Schuster, was the main investigator for this study. More precisely, I collaborated on conceptualization and writing of the research proposal, setting up the questionnaire and getting the approval of the institutional review board. I was responsible for data collection of dermatologic patients and coordinated the data collection of the other patient groups and controls. I was responsible for data handling and digitalization. I conducted all statistical analyses and wrote the draft of the manuscript, which I finalized with the valuable comments of my co-authors. Finally, I managed the submission process and took care of all required revisions.

## **1.2 Contribution to paper II**

This paper is based on the same data as paper I. I, Barbara Schuster, was the main investigator for this study. More precisely, I collaborated on conceptualization, writing of the research proposal, setting up the questionnaire and getting the approval of the institutional review board. I was responsible for data collection of dermatologic patients as well as for data handling and digitalization. I conducted all statistical analyses and wrote the draft of the manuscript, which I finalized with the valuable comments of my co-authors. Finally, I managed the submission process and took care of all required revisions.

## **1.3 Contribution to paper III**

I, Barbara Schuster, was involved in conceptualization and writing of the research proposal, setting up the questionnaire and getting the approval of the institutional review board. I was responsible for programming the online questionnaire as well as for data handling. I conducted all statistical analyses and wrote the draft of the manuscript, which I finalized with the valuable comments of my co-authors. Finally, I managed the submission process and, with the support of my co-authors, took care of all required revisions.

## 2. Introductory summary

### 2.1 Evaluating the mental burden of skin diseases

Skin diseases are common and very diverse. In the International Classification of Diseases (ICD)-11, more than 2000 skin diseases are recognized [1], ranging from acute infections to chronic congenital diseases and cancer. In spite of this diversity, most skin diseases have one thing in common which sets them apart from other diseases: as they affect the skin, which is the outermost layer of our body, they are often clearly visible to our social surrounding. And while a lot of skin diseases are not in fact infectious, subconscious evolutionary cognitions and a lack of education about skin diseases often lead to individuals with skin diseases being stigmatized and socially excluded [2]. In addition, due to these experiences and the existing beauty ideal of clear, flawless skin, people with skin diseases often suffer from low self-esteem and even body image disorders [3-6]. Overall, a large number of studies have shown that patients with skin diseases have a higher risk for mental comorbidities such as depression, anxiety, and addiction, than healthy individuals [7-10].

Over the last decades, the mental burden of skin diseases has received growing attention in dermatologic research, and a considerable body of data has been gathered by researchers all over the world [11-14]. However, previous research has predominantly taken a pathogenic perspective, assessing mental health by measuring mental comorbidities [7, 10, 15-17]. In contrast, it was already in 1948 when the World Health Organization (WHO) famously defined health as “a state of complete physical, mental and social well-being and not merely the absence of disease or infirmity” [18]. This implies that for the evaluation of mental health, assessing mental disorders and comorbidities is not sufficient, and that factors making people feel and stay well need to be considered as well [19]. As a consequence, patient reported outcome measures (PROMs) and especially the concept of health related quality of life (QoL) has received growing attention in medical research, starting in the 1970s [20]. However, tools for measuring health related QoL assess the patients' life circumstances using pre-defined criteria, such as mobility, security and health. While these certainly affect well-being, they rather measure antecedents of well-being than well-being itself [21].

Of course, medical research has not been the only discipline aiming to assess well-being. In fact, exploring well-being and its determinants is the main goal of positive psychology, a sub-discipline of psychology which “is the study of the conditions and processes that contribute to the flourishing or optimal functioning of people, groups, and institutions” [22]. In contrast to medical research, which has rather concentrated on the dimension of ill-being, positive psychology focuses on the dimension of well-being, exploring constructs such as happiness, flourishing, and flow. Therefore, positive-psychological concepts and measures contribute greatly to a holistic evaluation of well-being. However, with the exception of psychiatry,

positive-psychological measures have only rarely been examined in health care research so far (Macaskill, 2016).

One of the most researched constructs of positive psychology is happiness, a measure of hedonic well-being [23]. In contrast to eudaimonic well-being, which means living a meaningful life full of purpose, hedonic well-being means living a pleasant, enjoyable life [24]. While different definitions of happiness have been suggested in psychological literature, happiness is commonly operationalized as subjective well-being [25, 26], which is the “cognitive and affective evaluation of one’s life as a whole” [23]. The affective component of subjective well-being encompasses both positive affect (PA) and negative affect (NA), while the cognitive component of subjective well-being is defined as satisfaction with life (SWL) [27, 28]. Following this conceptualization, subjective well-being - and consequently happiness - is considered high if one experiences a lot of PA, little NA, and a general satisfaction with one’s life.

Studies exploring subjective well-being in patients with chronic diseases have been conducted before. The results suggest that patients with kidney diseases, stroke, and cancer, experience less subjective well-being than healthy individuals [29-31]. However, studies are rare and operationalizations differ, which makes it hard to compare results and facets of subjective well-being. In dermatology, only very few studies have explored single facets of well-being so far, yielding inconclusive results [32-35]. Thus, more studies were needed in order to clarify if happiness could be a useful tool to better understand the mental burden of skin diseases.

## **2.2 Happiness in dermatology**

The aim of this PhD project was to contribute to a more holistic understanding of the mental burden of skin diseases by introducing and evaluating happiness as a new PROM in dermatology. In order to achieve this goal, three research projects were conducted. While the aim of the first research project was to collect baseline data on happiness in dermatologic patients, the second and third projects aimed to further evaluate the informative value of happiness in dermatologic research by comparing the construct of happiness to both QoL and depression.

### **2.2.1 Paper I: Collecting baseline data on happiness in dermatology**

As the existing data on happiness in patients with skin diseases was very scarce, the aim of the first research project was to collect baseline data on happiness in patients with skin diseases and to put these results into perspective by comparing them to other patient groups and healthy controls. A cross-sectional study was conducted from December 2017 to April 2019. In total, 335 in- and outpatients at the Department of Dermatology and Allergy of the

Technical University of Munich with the diagnoses psoriasis, atopic eczema, nummular eczema, mastocytosis, and skin cancer (keratinocyte carcinoma and malignant melanoma) were recruited. The participants filled in a questionnaire consisting of validated scales measuring the three facets of subjective well-being according to Ed Diener [28]: PA and NA (measured both with the Positive and Negative Affect Schedule, PANAS [36], and the Scale of Positive and Negative Experience, SPANE [37]), and SWL (measured with the Satisfaction with Life Scale, SWLS [38]). Additional to this multidimensional operationalization of happiness, the participants were asked to indicate how happy they were overall on a scale from 0 (extremely unhappy) to 10 (extremely happy). The questionnaire furthermore contained questions on related constructs such as general and disease-related QoL, which were not further analyzed in this paper. In addition to the data collected via these patient questionnaires, diagnosis, disease severity and current treatment were documented by the treating physician.

In order to put the results of the dermatologic patients into perspective, we additionally recruited patients with human immunodeficiency virus (HIV, recruited at the Department of Inner Medicine II of the Technical University of Munich) and patients with chronic inflammatory bowel diseases (Crohn's disease and ulcerative colitis, recruited at Hospital Neumarkt in der Oberpfalz). These two patient groups were chosen for the following reasons: HIV is a chronic infectious disease which – if appropriately treated – does not go a long with physical impairment for the affected individual [39]. However, studies have shown that affected individuals often face stigmatization [40, 41]. In contrast, chronic inflammatory bowel diseases cause strong pain and lead to significant physical impairment for the affected individuals [42]. Both patients with HIV and chronic inflammatory bowel diseases filled in the same questionnaire as the dermatologic patients. Diagnosis, disease severity and treatment were documented by the treating physician. Last but not least, a group of healthy controls living in the greater Munich area was recruited with the help of a recruitment service (Testing Time, Zurich, Switzerland). The healthy controls filled in the same questions as the patient groups, but online.

Group comparisons of the z-standardized and age- and sex-adjusted means ( $m^a$ ) of the happiness variables showed that dermatologic patients reported significantly less PA ( $m^a = 3.18$  vs.  $3.49$ ;  $p = .001$ ) and judged themselves less happy than the healthy control group ( $m^a = 6.47$  vs.  $7.11$ ;  $p = .023$ ). In contrast, dermatologic patients were more satisfied with their lives than the control group ( $m^a = 5.13$  vs.  $4.83$ ;  $p = .043$ ). No significant difference was found in terms of NA ( $m^a = 1.99$  vs.  $1.93$ ;  $p = .418$ ). In comparison to the other patient groups, dermatologic patients reported less PA than patients with HIV ( $m^a = 3.18$  vs.  $3.48$ ;  $p = .016$ ), but comparable results compared with patients with chronic inflammatory bowel diseases ( $m^a = 3.18$  vs.  $2.97$ ;  $p = .08$ ). Differentiating the dermatologic diseases, we found that patients with psoriasis and atopic dermatitis reported lower PA (psoriasis:  $m^a = 3.20$ , atopic dermatitis:  $m^a = 3.01$ ) and judged themselves less happy (psoriasis:  $m^a = 5.85$ , atopic dermatitis:  $m^a =$

6.09) than healthy controls (all  $p < .05$ ), while patients with skin cancer reported elevated SWL compared to the control group ( $m^a = 5.53$  vs.  $4.81$ ;  $p = .003$ ).

The results indicate that patients with skin diseases are less happy than healthy individuals and patients with HIV. More precisely and most interestingly, they did not report more negative emotions, but less positive emotions. This finding underlines the importance of exploring positive markers of well-being such as happiness in this patient group, as otherwise the mental burden of skin diseases would be underrated. In addition, PA has been linked to several beneficial health outcomes such as better immune reactions, better cardiovascular health and even better skin barrier recovery [43-45]. Consequently, reduced levels of PA could contribute not only to the mental, but also to the physical burden of skin diseases. However, as this study was a cross-sectional study which does not allow to explore causality, more research is needed to verify this hypothesis. Comparing diagnoses, patients with psoriasis and atopic dermatitis seemed to be especially impaired in terms of happiness, which is in line with the large body of existing evidence pointing towards an especially large burden of these two chronic skin diseases [9, 17, 46-49]. In contrast, patients with skin cancer were more satisfied with their lives. While the existing study cannot provide an explanation for this finding, prior studies in patients with breast and colorectal cancer have shown that receiving a cancer diagnosis can trigger reevaluation processes [50], which might explain elevated SWL in skin cancer patients. Overall, these findings support the methodological approach of exploring happiness in patients with skin diseases and encourage further research in that matter.

The results of this study are published under the title “Happiness in Dermatology: a Holistic Evaluation of the Mental Burden of Skin Diseases” in the *Journal of the European Academy of Dermatology and Venereology* [51], where the paper was additionally chosen and presented as ‘Editor’s pick’.

### **2.2.2 Paper II: Differentiating happiness and (skin-related) quality of life**

Encouraged by the WHO’s definition of health as more than just the absence of disease [18], patients’ QoL has received growing attention in medical research, and more and more generic and disease-specific QoL questionnaires have been introduced [20]. While generic QoL questionnaires allow to measure and compare health-related QoL across different diagnoses and disciplines, disease-specific QoL questionnaires provide a more detailed and adapted analysis of the impact of different diseases on QoL. Today, health-related QoL is one of the most commonly used patient-reported outcomes both in clinical research and in practice [52] and is even listed as a relevant criterion for the evaluation of therapies both by the European Medicines Agency (EMA) and the American Food And Drug Administration (FDA) [53].

In dermatology, the Dermatology Quality of Life Index (DLQI) is the most commonly used quality of life questionnaire [54, 55]. It consists of 10 questions focusing on different aspects

of QoL impairment, e.g. the (in)ability to work or to do household chores due to the skin disease, or the impact of the skin disease on social and leisure life. The DLQI is frequently used in both clinical trials and practice in order to indicate the disease burden beyond the physical symptoms [54]. However, considering the WHO definition of health as a state of well-being [18], this raises the question whether the DLQI, which was originally developed as “disability index” [55], is sufficient in reflecting the patients’ disease burden, or whether indicators of well-being such as happiness can provide relevant information besides the DLQI. Thus, the aim of this second research project was to compare happiness to the most commonly used dermatologic QoL instrument, the DLQI. More precisely, the aim was to evaluate whether happiness provides information about the burden of skin diseases going beyond the DLQI, and to put the results into perspective by comparing both happiness and DLQI to a generic QoL instrument recommended by the WHO.

The data presented in this paper was collected in the same study as the data presented in Paper I of this PhD project. Besides the questions on happiness described in the previous chapter, the study questionnaire contained both a generic quality of life measure, namely the WHOQOL BREF (a 26-item short form of the more comprehensive WHOQOL) [56], and the DLQI [55] as a disease-specific QoL measure. For this paper, these results in terms of happiness and QoL were combined and compared. As the analyses in Paper I had shown that patients with different skin diseases show different patterns of well-being, and that patients with psoriasis and atopic dermatitis were especially impaired in terms of happiness [51], only patients with psoriasis and atopic dermatitis were included in this paper.

Linear regression models showed that the DLQI was strongly correlated with NA ( $\beta = -.49$ ;  $p < .001$ ), but not with PA ( $\beta = -.03$ ;  $p = .793$ ) nor SWL ( $\beta = -.2$ ;  $p = .173$ ) in patients with psoriasis and atopic dermatitis. Combining all three facets of happiness, the DLQI explained 23% of variance in happiness. When comparing both happiness and the DLQI to generic QoL, the DLQI explained 26% of variance in overall QoL and was associated only with the physical dimension of QoL ( $\beta = -.42$ ;  $p = .002$ ), but not with the psychological ( $\beta = -.02$ ;  $p = .189$ ), the social ( $\beta = .003$ ;  $p = .981$ ), nor the environmental domain ( $\beta = .01$ ;  $p = .905$ ). In contrast, all happiness variables together explained 70% of variance in generic QoL, with each facet of happiness contributing significantly to the model. Combining both overall happiness and the DLQI in one model predicting generic QoL, the amount of explained variance increased to 73%, with both the DLQI and happiness significantly adding to the model (DLQI:  $\beta = -.13$ ;  $p = .031$ ; happiness:  $\beta = .77$ ;  $p < .001$ ), indicating that both constructs carried unique information about overall QoL.

As a conclusion, we found that the DLQI predominantly measured NA and physical QoL impairment, which is in line with its original purpose of being a disability index. However, the DLQI alone only provided a limited evaluation of QoL as recommended by the WHO. Adding

happiness as a new and very subjective patient reported outcome provides additional information on the mental burden of skin diseases not yet covered by the DLQI and contributes to a more comprehensive assessment of QoL without the need for an additional generic and time-consuming quality of life assessment.

The results of this project have been accepted for publication under the title “Adding happiness to complement the Dermatology Quality of Life Index in Psoriasis and Atopic Dermatitis Healthcare” in the *European Journal of Dermatology* [57].

### **2.2.3 Paper III: Differentiating happiness and depression**

Prior research on the mental burden of skin diseases has mostly taken a pathogenic perspective, assessing mental comorbidities such as depression, anxiety or addiction as indicators of mental health. Especially for psoriasis, a large number of studies have demonstrated a higher risk of mental comorbidities [9, 10, 46, 48]. These findings are one of the reasons why psoriasis was defined as a “serious non-communicable disease” by the World Health Assembly in 2014 [58].

However, bearing the WHO’s definition of health in mind [18], assessing mental comorbidities only measures one side of the mental burden of skin diseases. In order to achieve a more holistic understanding, researchers and the WHO alike are demanding for a more holistic evaluation of the mental burden of psoriasis, which incorporates measures of well-being [7, 59, 60]. However, in psychological research there is still an ongoing debate whether depression and happiness are indeed two different constructs [61, 62]. This debate is especially relevant as depression, amongst other symptoms, is characterized by the inability to feel happy [63]. This raises the question whether assessing happiness in patients with skin diseases does actually provide new insights of the mental burden of skin diseases, or whether it just reflects the findings of a high risk for depression in this population. Therefore, the aim of this study was to simultaneously assess and compare happiness and depression in a large and diverse sample of patients with psoriasis and to explore determinants of both constructs.

For this project, a cross-sectional online survey was conducted from March to June 2019. The link to an online questionnaire was shared on Germany’s largest patients’ information website about psoriasis ([www.psoriasis-netz.de](http://www.psoriasis-netz.de)) as well as on the Facebook pages of several psoriasis associated organizations or campaigns (e.g. Farbenhaut, “bitte berühren” of the Professional Association of German Dermatologists). The questionnaires contained validated scales measuring subjective well-being (SPANE, SWLS) [37, 38], as well as a validated screening questionnaire for depression (WHO-5 Well-being index) [64]. Additionally, current and general disease severity were assessed using self-evaluation (mild, moderate, severe) and data on treatment status and years since the first diagnosis were collected.

In total, the data of 722 participants with psoriasis were analyzed. Controlling for sex and age, we found that the participants reported lower PA ( $p = .003$ ) and lower SWL ( $p < .001$ ;

except participants aged 65+ years) than German reference populations (norm data or in case of PA and NA, data of the German validation sample), but not more NA ( $p = .767$ ). 40.3% of participants were screened positive for depression. Using exploratory factor analysis, PA, NA, SWL and depression were confirmed as four distinct constructs. General disease severity was identified as main risk factor for low happiness and depression. In contrast, participants who were currently in a phase of improvement of skin condition reported higher happiness and less depressive tendencies. Similarly, the longer the participants had been living with their diagnosis the more happy and less depressed they were.

As a conclusion, the results of this study not only confirmed the findings of low PA in patients with psoriasis (as presented in Paper I of this PhD project) in a large and diverse sample of patients with psoriasis. They also supported the methodological approach of differentiating happiness in depression in patients with psoriasis.

The results of this project are published under the title “Happiness and depression in psoriasis: a cross-sectional study in Germany” in the Journal *Quality of Life Research* [57].

### **2.3 Conclusion**

As a conclusion, this PhD thesis showed that assessing happiness contributes to a more holistic evaluation of the mental burden of skin diseases which goes beyond mental comorbidities and QoL. The results indicate that patients with skin diseases, and especially with psoriasis and atopic dermatitis, seem to experience less PA than healthy individuals, but not necessarily more NA. As PA has only rarely been studied in patients with skin diseases so far, this part of the mental burden of skin diseases has been overlooked so far. The finding of low PA is especially interesting as PA has been shown to directly and indirectly affect physical health by influencing our endocrine system, our immune responses, and our health behavior [65, 66]. Future research as well as clinical practice should start to incorporate measures of well-being such as happiness in order to better understand the patients' needs and to provide personalized medical care. Additionally, future research should explore ways of increasing PA in patients with skin diseases, as this could not only lower the mental burden of skin diseases, but potentially contribute to better physical health and maybe even improvement of skin condition.



### 3. Paper I

#### **Happiness in Dermatology: a Holistic Evaluation of the Mental Burden of Skin Diseases**

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## ORIGINAL ARTICLE

## Happiness in dermatology: a holistic evaluation of the mental burden of skin diseases

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### Abstract

**Background** According to the World Health Organization, mental health is a state of well-being and not merely the absence of disease. However, studies exploring subjective well-being in patients with skin diseases are very rare.**Objectives** To assess subjective well-being, i.e. 'happiness', in patients with different skin diseases and to compare them to other patient groups and healthy controls.**Methods** A cross-sectional study was conducted from 12/2017 to 04/2019. Patients receiving in- or outpatient care for psoriasis, atopic eczema, nummular eczema, mastocytosis, skin cancer (malignant melanoma and keratinocyte carcinoma), human immunodeficiency virus (HIV) or chronic inflammatory bowel diseases (Crohn's disease and ulcerative colitis) were recruited at two hospitals in Bavaria, Germany. Healthy individuals living in or near Munich served as a control group. All participants filled in a questionnaire assessing happiness, measured as positive affect (PA), negative affect and satisfaction with life (SWL; together representing subjective well-being) and a heuristic evaluation of one's own happiness.**Results** Data from 229 dermatologic patients (53.3 ± 18.5 years, 48% women), 49 patients with inflammatory bowel diseases (48.9 ± 18.7 years, 43% women), 49 patients with HIV (46 ± 10.1 years, 10% women) and 106 healthy controls (38.4 ± 13.4 years, 49% women) were analysed. Compared to the controls, dermatologic patients reported lower heuristic happiness ( $P = 0.023$ ) and PA ( $P = 0.001$ ) but higher SWL ( $P = 0.043$ ). Patients with psoriasis and atopic eczema reported the lowest happiness, as they reported significantly lower PA ( $P = 0.032$  and  $P < 0.001$ ) and heuristic happiness ( $P = 0.002$  and  $P = 0.015$ ) than the control group. Patients with skin cancer reported higher SWL than the control group ( $P = 0.003$ ). Dermatologic patients reported lower happiness than patients with HIV but reported greater happiness than patients with IBD.**Conclusions** Dermatologic patients experience lower levels of happiness, especially PA, compared to healthy controls. As PA is linked to desirable health outcomes, targeting PA could be a promising holistic approach for the treatment of skin diseases.

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### Conflict of interest

The authors declare no conflict of interest.

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### Introduction

Patients with skin diseases have a higher risk for mental comorbidities such as depression, anxiety and addiction.<sup>1–4</sup> While thereis a growing body of research on this topic, it is striking that almost all studies exploring the mental burden of skin diseases focused on assessing mental disorders.<sup>2,5–7</sup> However, considering the World Health Organization's (WHO) definition of health as a 'state of well-being and not merely the absence of disease',<sup>8</sup>

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such studies are only able to provide a partial assessment of the full mental burden of skin diseases. Studies measuring mental health in terms of subjective well-being are almost absent in dermatologic research so far.

While there are different definitions, happiness is often defined as subjective well-being,<sup>9</sup> which represents 'a person's cognitive and affective evaluations of his or her life as a whole'.<sup>10</sup> Following this definition, subjective well-being is considered high if one experiences plenty of positive emotions, few negative emotions and is generally satisfied with one's life.<sup>10,11</sup>

Integrating well-being seems a promising approach, especially in dermatology, as studies have found significant relationships between psychosocial factors, mostly stress-related, on the occurrence, severity and progression of skin diseases and that psychological factors can play a role in the treatment of the diseases.<sup>12–14</sup> For example, a prospective study among 62 psoriasis patients over the course of 6 months showed that stressful periods were related to higher disease severity 4 weeks later, but only in patients who tend to worry or scratch.<sup>15</sup> Similarly, other studies showed that psoriasis patients who worry a lot are less responsive to psoralen and ultraviolet A (PUVA) therapy<sup>16</sup> and that listening to meditation audiotapes during phototherapy (UVB and PUVA) leads to better treatment results in patients with psoriasis.<sup>17</sup> In addition, positive affect (PA) has been shown to be associated with several desirable health outcomes, ranging from improved health behaviour and better cardiovascular health to better immune responses, quicker wound healing and reduced pain.<sup>18–20</sup> Apart from these beneficial outcomes, the well-being of patients is a valuable aim in itself. Thus, exploring well-being among affected patients utilizing diverse measures could greatly contribute to a better understanding and treatment of skin diseases.

However, in dermatologic research, happiness and subjective well-being have been widely neglected as outcome variables with only very few exceptions. Only one study measured PA in patients with psoriasis,<sup>21</sup> while three studies looked at negative affect (NA) and life satisfaction as components of well-being.<sup>22–24</sup> While these studies underline the importance of happiness as a patient-reported outcome, their informative value is limited as they measured only components of happiness, were restricted to only one patient group, lacked a control group and used self-reported scores of disease severity.

To gain a more balanced and valid view on the mental burden of skin diseases, this explorative study sought to assess happiness and subjective well-being holistically, using PA, NA, satisfaction with life (SWL) and a heuristic measure of happiness. We assessed patients with different skin diseases and compared them to patients with chronic non-dermatologic diseases as well as healthy controls. In addition, disease severity was documented by the attending physician using validated indices.

## Materials and methods

This cross-sectional study was conducted from December 2017 to April 2019. The dermatologic diagnoses examined were psoriasis, atopic eczema, nummular eczema, mastocytosis and skin cancer (malignant melanoma and keratinocyte carcinoma). For comparison purposes, patients with chronic inflammatory bowel diseases (IBD: ulcerative colitis and Crohn's disease), which can be very physically demanding due to strong pain during flares,<sup>25</sup> patients with infection with human immunodeficiency virus (HIV) who due to antiretroviral therapy are mainly physically unimpaired<sup>26</sup> but may have been experiencing stigmatization<sup>27,28</sup> and a healthy control group were included.

Patients with skin diseases were recruited from the Department of Dermatology and Allergy and patients with HIV from the interdisciplinary HIV centre at University hospital of the Technical University of Munich, while patients with IBD were recruited from the Department of Internal Medicine II, Hospital Neumarkt in der Oberpfalz. Only patients aged 18 years or older, suffering from one of the examined diagnoses, receiving in- or outpatient medical care at the respective clinics were eligible to participate in this study. Patients meeting these criteria were asked by the attending physician or assistant to fill in a paper-based questionnaire. Patients unable to fill in the German paper-based questionnaire were excluded. Written informed consent was obtained from all patients prior to study inclusion.

Healthy individuals, who served as a control group, were recruited with the help of a recruitment service (TestingTime, Zurich, Switzerland). Only individuals aged 18 years or older living in the greater Munich area and not suffering from one of the examined diseases were eligible to participate. Participants in the control group filled in an online version of the paper-based questionnaire used for the patient groups. The TestingTime recruiting team was responsible for the recruitment and the financial compensation of the participants (3€ per person) but was not involved in the data collection. Only the participants of the control group (as recruited by TestingTime) received a financial compensation.

This study was conducted in accordance with the Declaration of Helsinki and was approved by the local ethics committee of the Technical University of Munich (reference number 424/17S).

## Questionnaire

The questionnaire consisted of validated scales measuring happiness and subjective well-being. For the patients, the attending physician additionally documented diagnoses and, when clinical scores were available, disease severity based on the following scores: Psoriasis Area and Severity Index for psoriasis, Scoring Atopic Dermatitis for atopic dermatitis, Mayo score & Crohn's disease activity index for IBD and Centers of Disease Control and Prevention (CDC) stage for HIV. The respondents were further asked to report any comorbidities in the questionnaire.

Positive affect and NA were measured using the German version of the Positive and Negative Affect Schedule (PANAS),<sup>29,30</sup> which consists of 20 adjectives (10 representing PA, e.g. 'excited' and 'enthusiastic', and 10 representing NA, e.g. 'distressed' and 'upset'), each rated on a 5-point scale ranging from 1 ('very slightly or not at all') to 5 ('extremely'). The PANAS is the most commonly used instrument for measuring affect.<sup>31,32</sup> The two distinct subscales, PA and NA, showed good reliability in this sample with Cronbach's alphas of 0.92 and 0.86, respectively. As the PANAS measures rather specific emotions,<sup>9</sup> we additionally used the German version of the Scale of Positive and Negative Experience (SPANE), which measures PA and NA in a more comprehensive way using 12 more general items (e.g., 'pleasant' and 'unpleasant').<sup>32</sup> Both subscales of SPANE showed good reliability in this sample with Cronbach's alphas of 0.94 for PA and 0.88 for NA. For both the PANAS and SPANE, the items of each subscale were averaged to form an index if at least 80% of items were answered validly.

Satisfaction with life was measured using the SWL Scale (SWLS),<sup>33</sup> which consists of five items (e.g. 'in most ways my life is close to my ideal'), each rated on a seven-point scale ranging from 1 ('strongly disagree') to 7 ('strongly agree'). The scale showed good reliability in this sample ( $\alpha = 0.88$ ), and the items were averaged to form an index if at least 80% of items were answered validly.

In addition to this theory-based operationalization of happiness, we included a measure of *heuristic happiness* in the form of a single question derived from the European Social Survey<sup>34</sup>: 'Taking all things together, how happy would you say you are?' Respondents could give their answer on an 11-point scale ranging from 0 ('extremely unhappy') to 10 ('extremely happy').

### Statistics

In order to identify and eliminate potential errors in manual data digitalization, data collected via paper-based questionnaires were digitalized twice using Epi Info™ (CDC, Atlanta, GA, USA) and any discrepancies between the resulting datasets were sorted out. Patients suffering from more than one of the examined diseases were excluded from all further analyses to avoid ambiguous results. Differences in age and gender between the groups were analysed using a Student's *t*-test and a chi-squared test. For the analysis of PA and NA, we focused on the PANAS, because it is the most commonly used tool for measuring affect.<sup>31,32</sup> In case of divergent results, we also reported the results obtained with SPANE for a more comprehensive evaluation of subjective well-being (see also Table S1, Supporting Information for the detailed results of the SPANE). Analyses of Covariance (ANCOVAs) were used to compare the groups of participants while controlling for gender and age. Age- and gender-adjusted means ( $m_a$ ) and partial eta squared ( $\eta^2$ ) are reported. Planned contrasts were calculated to reveal differences between the groups. To increase comparability, z-standardized data are presented when

appropriate. To analyse the relationship between disease severity and happiness, z-standardized clinical scores were entered into linear regression models, with the different happiness measures as outcomes and age and gender as covariates. Standardized regression coefficients ( $\beta$ ) are reported.

A main symptom of clinical depression is the inability to feel happy.<sup>35</sup> To rule out potential confounding, we conducted a sensitivity analysis and excluded all participants suffering from depression (four in total: one psoriasis, one atopic eczema, one mastocytosis and one ulcerative colitis) as assessed using the provided information on comorbidities in the questionnaire. As the exclusion did not have significant influence on the results, we reported the final results including the depressed participants.

The level of significance was set at  $\alpha = 0.05$  for all analyses. All statistical analyses were conducted using IBM SPSS Statistics Version 24 (IBM Corporation, Armonk, NY, USA).

### Results

In total, 335 patients and 106 controls were recruited. Of those, eight patients affected by more than one of the examined diseases were excluded. Of the remaining 327 patients, 229 were dermatologic patients (52 psoriasis, 50 atopic eczema, 24 nummular eczema, 53 mastocytosis and 50 skin cancer), 49 patients had been diagnosed with IBD (28 Crohn's disease and 21 ulcerative colitis) and 49 patients with HIV. The examined groups differed significantly in terms of gender ( $P < 0.001$ ) and age ( $P < 0.001$ ; Tables 1 and 2).

Unadjusted means, medians, quartiles, minima and maxima of the examined happiness variables are displayed in Fig. 1.

#### Comparing dermatologic patients to patients with HIV, IBD and healthy controls

Taking all dermatologic diseases together, we found significant effects of group membership (patient groups and controls) on heuristic happiness ( $\eta^2 = 0.028$ ,  $P = 0.009$ ) and PA ( $\eta^2 = 0.051$ ,  $P < 0.001$ ), but not on NA ( $\eta^2 = 0.004$ ,  $P = 0.612$ ) and SWL ( $\eta^2 = 0.010$ ,  $P = 0.223$ ). Compared to the control group, dermatologic patients reported lower heuristic happiness ( $P = 0.023$ ) and PA ( $P = 0.001$ ) but higher SWL ( $P = 0.043$ ; Table 3 and Fig. 2). No significant difference was found for NA ( $P = 0.418$ ). Compared to the other patient groups, dermatologic patients reported significantly lower PA than patients with HIV ( $P = 0.016$ ) and higher PA than patients with IBD, but this difference was not significant ( $P = 0.08$ ). Dermatologic patients did not significantly differ from HIV and IBD patients regarding NA, SWL and heuristic happiness.

#### Comparing patients with different dermatologic diseases to healthy controls

When looking separately at the different dermatologic diseases compared to controls, we found effects of group membership on heuristic happiness ( $\eta^2 = 0.050$ ,  $P = 0.006$ ) and

**Table 1** Participants' characteristics

	Dermatology total <i>n</i> = 229†	Inflammatory bowel diseases <i>n</i> = 49‡	HIV <i>n</i> = 49§	Control group <i>n</i> = 106
<b>Age</b>				
Mean + SD	53.3 ± 18.4	48.9 ± 18.7	46 ± 10.1	38.4 ± 13.4
18–29 years	29 (13%)	9 (18%)	2 (4%)	32 (30%)
30–44 years	43 (19%)	12 (25%)	20 (41%)	40 (38%)
45–64 years	80 (35%)	19 (39%)	25 (51%)	30 (28%)
65+ years	75 (33%)	9 (18%)	2 (4%)	4 (4%)
<b>Gender</b>				
Men	119 (52%)	27 (57%)	44 (90%)	54 (51%)
Women	109 (48%)	20 (43%)	5 (10%)	52 (49%)
<b>Disease severity¶</b>				
Mean + SD/%	PASI: 12.7 ± 9.6 SCORAD: 42.3 ± 19.9	CDAI: 102.7 ± 108.3 Mayo: 4.5 ± 3.5	CDC stage A: 19 (46%) CDC stage B: 7 (17%) CDC stage C: 15 (37%)	–

†Two missing cases for age, one missing case for gender, ‡Two missing cases for gender, §Eight missing cases for CDC stage, ¶Data on disease severity are only available for patients with psoriasis (Psoriasis Area and Severity Index, PASI; score ranging from 0 to 72; *n* = 48), atopic eczema (Scoring Atopic Dermatitis, SCORAD; score ranging from 0 to 103; *n* = 45), Crohn's disease (Crohn's Disease Activity Index, CDAI; score ranging from 0 to 600; *n* = 28), ulcerative colitis (Mayo score; score ranging from 0 to 12; *n* = 19) and human immunodeficiency virus (HIV; Centers for Disease Control and Prevention CDC stages; *n* = 41).

PA ( $\eta^2 = 0.045$ ,  $P = 0.011$ ), but not on NA ( $\eta^2 = 0.025$ ,  $P = 0.160$ ) and SWL ( $\eta^2 = 0.032$ ,  $P = 0.063$ ). Specifically, we found significantly lower heuristic happiness and PA in patients with psoriasis and atopic eczema compared to the control group (happiness:  $P = 0.002$  and  $P = 0.015$ ; PA:  $P = 0.032$  and  $P < 0.000$ ; Table 3 and Fig. 2). Patients with mastocytosis and nummular eczema reported lower PA compared to the controls, but the differences did not reach significance ( $P = 0.050$  and  $P = 0.081$ ). Patients with skin cancer reported significantly higher SWL than the control group ( $P = 0.003$ ). Regarding patients with mastocytosis, elevated SWL did not reach significance in comparison with

the controls ( $P = 0.074$ ). For NA, there were no differences between patients with skin diseases and the control group.

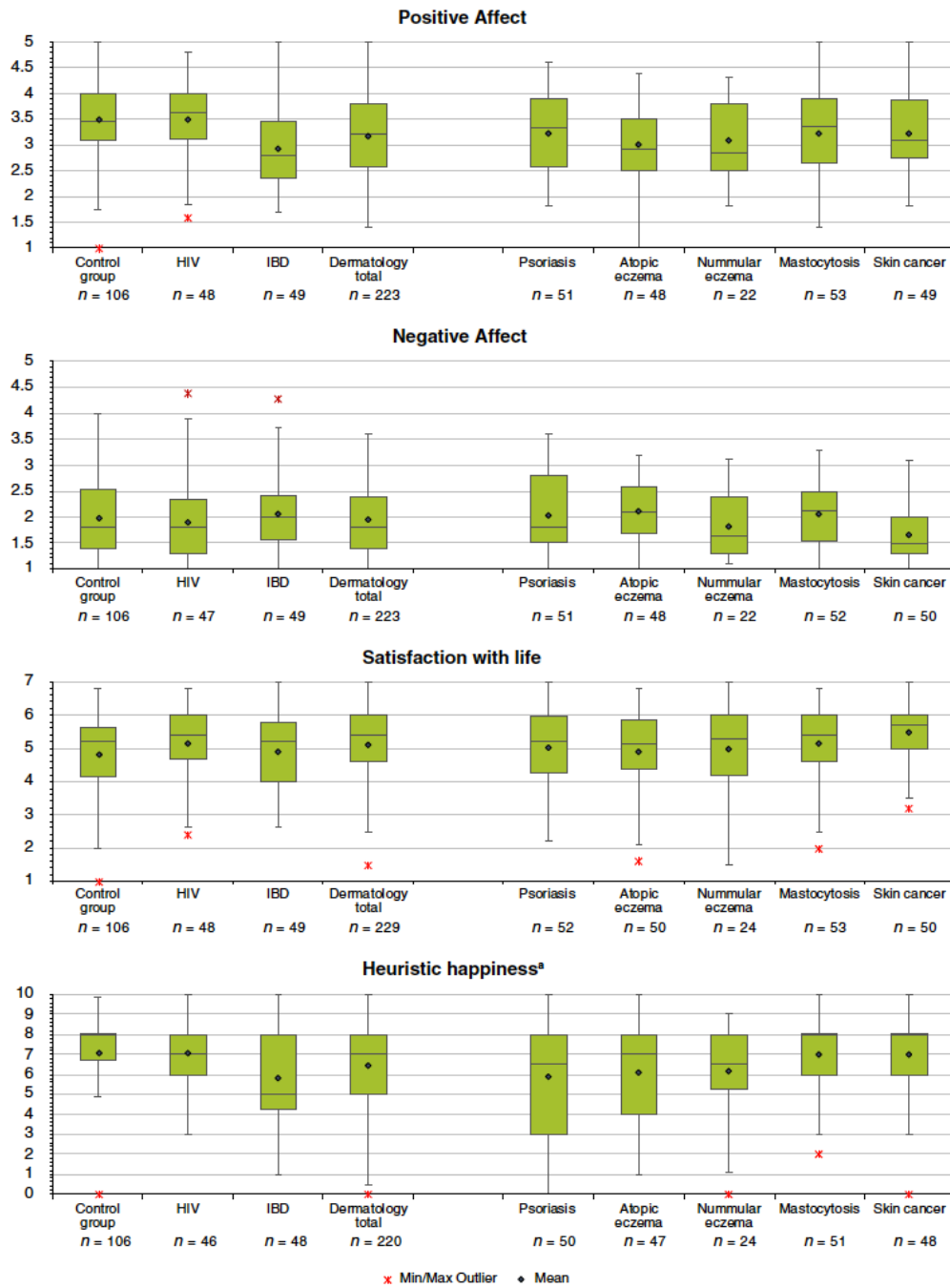
#### The role of disease severity

Taking all diagnoses with clinical scores for disease severity available together (psoriasis, atopic dermatitis, HIV and IBD), we found that higher disease severity was associated with lower PA ( $\beta = -0.20$ ,  $P = 0.007$ ). When differentiating between the diseases, we found a similar association with disease severity for patients with psoriasis ( $\beta = -0.33$ ,  $P = 0.041$ ) and IBD ( $\beta = -0.49$ ,  $P = 0.002$ ), but not for patients with atopic eczema ( $\beta = 0.71$ ,  $P = 0.694$ ) and HIV ( $\beta = -0.36$ ,  $P = 0.723$ ). Disease

**Table 2** Patient characteristics of dermatologic patients

	Psoriasis <i>n</i> = 52†	Atopic eczema <i>n</i> = 50	Nummular eczema <i>n</i> = 24	Mastocytosis <i>n</i> = 53‡	Skin cancer <i>n</i> = 50
<b>Age</b>					
Mean + SD	48.7 ± 16.8	46 ± 21.2	60.8 ± 13.3	47.1 ± 13.7	68 ± 13.1
18–29 years	10 (20%)	14 (28%)	0	5 (10%)	0
30–44 years	9 (18%)	12 (24%)	4 (17%)	17 (33%)	1 (2%)
45–64 years	21 (41%)	12 (24%)	5 (21%)	21 (40%)	21 (42%)
65+ years	11 (22%)	12 (24%)	15 (63%)	9 (17%)	28 (56%)
<b>Gender</b>					
Men	31 (62%)	26 (52%)	14 (58%)	15 (29%)	32 (64%)
Women	20 (39%)	24 (48%)	10 (42%)	37 (71%)	18 (36%)
<b>Disease severity§</b>					
Mean + SD	12.7 ± 9.6	42.3 ± 19.9	–	–	–

†One missing case for age, ‡One missing case for age and gender, §Data on disease severity are only available for patients with psoriasis (Psoriasis Area and Severity Index, PASI; score ranging from 0 to 72; *n* = 48) and atopic eczema (Scoring Atopic Dermatitis, SCORAD; score ranging from 0 to 103; *n* = 45).



**Figure 1** Unadjusted means, medians, quartiles and minima/maxima of positive affect, negative affect, life satisfaction and heuristic happiness. <sup>a</sup>Heuristic evaluation of own happiness on a scale ranging from 0 to 10.

**Table 3** Descriptive statistics for the happiness measures (patient groups and healthy controls)

	Positive affect		Negative affect		Satisfaction with life		Heuristic happiness†	
	<i>n</i> , <i>m</i> ± SD	<i>m</i> <sub>a</sub>	<i>n</i> , <i>m</i> ± SD	<i>m</i> <sub>a</sub>	<i>n</i> , <i>m</i> ± SD	<i>m</i> <sub>a</sub>	<i>n</i> , <i>m</i> ± SD	<i>m</i> <sub>a</sub>
<b>All diseases</b>								
Dermatology total	223, 3.17 ± 0.76	3.18	223, 1.96 ± 0.66	1.99	229, 5.12 ± 1.17	5.13	220, 6.48 ± 2.34	6.47
Inflammatory bowel diseases	49, 2.95 ± 0.76	2.97	49, 2.08 ± 0.67	2.08	49, 4.93 ± 1.16	4.99	48, 5.81 ± 2.21	5.89
Human immunodeficiency virus	48, 3.51 ± 0.71	3.48	47, 1.91 ± 0.72	1.95	48, 5.14 ± 1.10	5.11	46, 7.09 ± 1.64	6.98
Healthy controls	106, 3.51 ± 0.71	3.48‡	106, 2.00 ± 0.71	1.93‡	106, 4.82 ± 1.26	4.83‡	106, 7.08 ± 2.06	7.11‡
<b>Dermatologic diseases</b>								
Psoriasis	51, 3.22 ± 0.79	3.20	51, 2.05 ± 0.77	2.08	52, 5.02 ± 1.14	5.02	50, 5.92 ± 2.81	5.85
Atopic eczema	48, 3.02 ± 0.66	3.01	48, 2.11 ± 0.60	2.09	50, 4.90 ± 1.35	4.89	47, 6.11 ± 2.35	6.09
Nummular eczema	22, 3.11 ± 0.73	3.16	22, 1.83 ± 0.63	1.89	24, 4.98 ± 1.45	4.99	24, 6.21 ± 2.40	6.20
Mastocytosis	53, 3.24 ± 0.84	3.22	52, 2.07 ± 0.60	2.06	53, 5.15 ± 1.10	5.18	51, 7.00 ± 1.80	7.07
Skin cancer	49, 3.24 ± 0.75	3.30	50, 1.66 ± 0.56	1.76	50, 5.50 ± 0.87	5.53	48, 7.02 ± 2.13	7.00
Healthy controls	106, 3.51 ± 0.71	3.48‡	106, 2.00 ± 0.71	1.96‡	106, 4.82 ± 1.26	4.81‡	106, 7.08 ± 2.06	7.07‡

†Evaluation of one's own happiness on a scale ranging from 0 to 10, ‡Due to the adjustment procedure, *m*<sub>a</sub> of healthy controls differs depending on which groups they were compared to (all diseases or dermatologic diseases only). Unadjusted means, standard deviation and age- and gender-adjusted means of the different happiness measures in the patient groups and the healthy controls. *m*, mean, *m*<sub>a</sub>, age- and gender-adjusted means; SD, standard deviation.

severity was not associated with any of the other happiness measures.

#### Alternative measurement of affect with SPANE

When measuring PA and NA with SPANE (Table S1, Supporting Information), some results differed from those obtained with the PANAS. In contrast to the findings reported above, the difference between dermatologic patients (*m*<sub>a</sub><sup>SPANE</sup> = 3.57) and healthy controls (*m*<sub>a</sub><sup>SPANE</sup> = 3.76) regarding PA<sup>SPANE</sup> did not reach significance (*P* = 0.063). Similarly, only patients with atopic eczema (*m*<sub>a</sub><sup>SPANE</sup> = 3.28) reported significantly lower PA<sup>SPANE</sup> than the healthy controls (*m*<sub>a</sub><sup>SPANE</sup> = 3.74, *P* = 0.001), but not patients with psoriasis (*m*<sub>a</sub><sup>SPANE</sup> = 3.53, *P* = 0.16). Regarding NA<sup>SPANE</sup>, patients with atopic eczema (*m*<sub>a</sub><sup>SPANE</sup> = 2.55) reported significantly higher values than the control group (*m*<sub>a</sub><sup>SPANE</sup> = 2.21, *P* = 0.019), which was not observed when measuring NA with PANAS.

#### Discussion

This study sought to extend previous research on the mental burden of skin diseases, which so far has almost exclusively focused on mental comorbidities related to skin diseases, by taking a holistic approach and measuring well-being and happiness. We found that dermatologic patients reported lower levels of happiness than participants of a control group both when using a heuristic and a theoretical operationalization of happiness. This finding is in accordance with previous studies that also reported reduced subjective well-being in patients with chronic diseases.<sup>36,37</sup> Dermatologic patients further reported lower levels of happiness than patients with HIV but scored higher than patients with chronic inflammatory bowel diseases.

#### Positive and negative affect

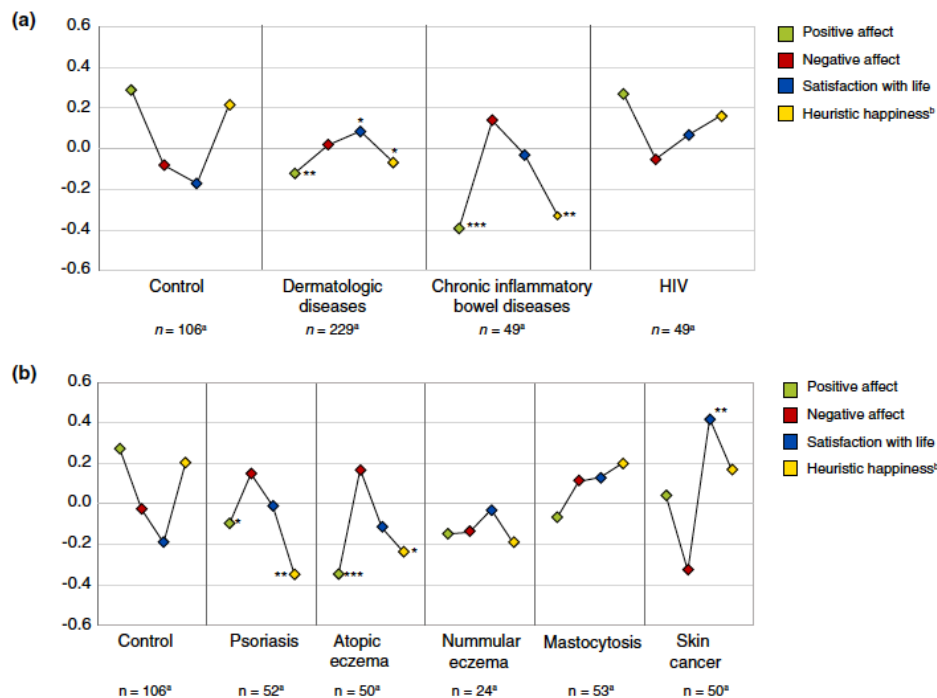
Patients with skin diseases reported lower PA as measured by the PANAS, but no differences emerged regarding NA compared to healthy controls. This finding is especially interesting, as PA has been shown to be associated with several desirable health outcomes, such as improved health behaviour, quicker wound healing, better cardiovascular health and better immune responses.<sup>18–20</sup> Thus, increasing PA in patients with skin diseases could not only improve well-being, but might even lead to a better prognosis.<sup>38</sup>

There is large evidence on interventions to promote PA.<sup>39,40</sup> Interventions that have been found to be highly effective are, for example, a gratitude journal or interventions focusing on meaning or strength use in daily life.<sup>39</sup> While these and further interventions have been positively evaluated, they are yet to be adapted to dermatologic patients.

It has to be added that the scale used to measure PA in this study (PANAS) focused on high-arousal positive emotions (e.g., 'excited' and 'enthusiastic').<sup>31</sup> When measuring affect with more general items (e.g. 'positive and 'pleasant'), the differences regarding PA between dermatologic patients and healthy controls were less pronounced and not significant (*P* = 0.063). Thus, the effect of skin diseases on PA might at least to some degree be moderated by arousal.

#### Comparing dermatologic diseases

Taking all findings together, patients with psoriasis and atopic eczema seemed to have the lowest levels of happiness, as they reported lower PA and lower overall happiness compared to participants of the control group. This corroborates previous research that has shown that psoriasis and atopic eczema are associated with several mental disorders, such as depression,



**Figure 2** Patterns of well-being in (a) all participants (b) dermatologic patients. Z-standardized, and age- and gender-adjusted means are shown. Asterisks indicate significant differences compared to the control group: \* $P < 0.05$ , \*\* $P < 0.01$ , \*\*\* $P < 0.001$  <sup>a</sup>Case numbers for different variables slightly vary due to missing data and <sup>b</sup>heuristic evaluation of own happiness on a scale ranging from 0 to 10.

addiction and even suicidal ideation.<sup>4,5,7</sup> In contrast, patients with skin cancer seemed to be the least unhappy in this sample; this finding is in accordance with a study conducted by Wikman and colleagues who found that patients with diabetes and cancer had higher levels of happiness than patients with other diseases including stroke and lung diseases.<sup>37</sup>

**Elevated satisfaction with life in patients with skin cancer**

Surprisingly, skin cancer patients reported significantly higher levels of SWL than all other groups. Even though skin cancer is a potentially fatal diagnosis, the prognosis is usually favourable if tumours are detected early, especially, but not only, in cases of keratinocyte carcinoma.<sup>41-43</sup> Accordingly, patients might be relieved after treatment and, consequently, feel more satisfied with their lives. Another explanation for their high levels of life satisfaction could be that the emotionally charged diagnosis of cancer may trigger re-evaluation processes in the affected patients. This has been observed for patients diagnosed with other types of cancer such as breast and colorectal cancer.<sup>44</sup> However, re-evaluation processes in patients diagnosed with skin cancer, which has a very low mortality rate compared to breast and

colorectal cancer, might imply different kinds of emotions and cognitions which remain to be tested.

The group of skin cancer patients examined in this study was quite heterogeneous and patients with melanoma and keratinocyte carcinoma (including its different types) and different tumour stages were not differentiated due to sample size restrictions. As both types of skin cancer and tumour stage greatly impact treatment and prognosis,<sup>42,43</sup> it seems probable that these variables might influence emotional reactions and well-being as well. For verification, more detailed studies with larger sample sizes allowing a closer differentiation between melanoma and keratinocyte carcinoma and different tumour stages are necessary.

**The role of dermatologic disease severity**

Higher disease severity was associated with less PA in patients with psoriasis but not in patients with atopic eczema. Thus, patients with more severe forms of psoriasis tend to experience less PA than those with milder forms, whereas patients with atopic eczema are affected in the same way regardless of disease severity. On the one hand, this could be due to differences in the symptoms caused by psoriasis and atopic eczema. For example, Evers and colleagues found that patients with atopic eczema



reported higher levels of itch than patients with psoriasis.<sup>45</sup> However, they also found that itch hardly contributed to levels of distress in both patient groups. In contrast, they identified fatigue as an important factor causing psychological distress, which was more pronounced in patients with atopic eczema than psoriasis. This might explain why milder forms of atopic eczema might be more mentally straining than milder forms of psoriasis. On the other hand, Bahmer and colleagues found that patients with psoriasis and atopic eczema differed regarding personality and character traits, with patients with psoriasis being less ambitious and narcissistic and patients with atopic eczema being less self-critical than the population average.<sup>22</sup> In addition, psoriasis has been linked to alexithymia,<sup>46,47</sup> which involves difficulties in recognizing and describing emotions. Such psychometric differences might lead to different ways of perceiving and coping with the respective skin disease,<sup>48</sup> which could explain the different effects of disease severity on PA. Finally, Schut and colleagues<sup>49</sup> found that in patients with atopic eczema, illness perception was significantly associated with self-assessed, but not physician-assessed physical health. Thus, the perception of disease severity of AD seems to differ between patients and physicians, which could explain why we did not find an association between objectively assessed disease severity and PA in those patients.

#### Strengths and limitations

This study is one of the very few studies<sup>21–24</sup> to examine happiness and subjective well-being in the context of dermatology. As there is no uniform operationalization of happiness, this study used a theory-based and a heuristic approach as well as two different instruments for measuring affect to increase validity. Furthermore, due to the recruitment of a control group living in the same area as the dermatologic patients and age- and gender-adjusted analyses, a valid comparison with healthy controls was possible that did not rely on unadjusted norm data as was the case in other studies.<sup>23</sup> A comparison with two different patient groups further enhanced the informative value of the results obtained in this study. In addition, a sensitivity analysis was conducted in order to rule out depression as a potential confounder.

Still, this study is subject to several limitations. First, only dermatologic patients receiving in- or outpatient care at a university hospital were included. As a result, severe cases might have been over-represented in this sample. While patients with HIV were recruited from the same hospital, IBD patients were recruited from a municipal hospital located in a more rural area. Thus, regional differences could have confounded the results of the comparison with this patient group. Furthermore, patients with mastocytosis were contacted by mail. Again, this might have affected the results for this group. Participants of the control group filled in an online version of the paper-based questionnaire used in the

patient groups. These different modalities of data collection could also have influenced the results in this study. Furthermore, depression was assessed using a question asking for current comorbidities, which implies that the respective diseases, in this case depression, have actually been diagnosed by a physician. Depressive symptoms or depressive tendencies were not assessed. Together with the fact that patients with acute clinical depression are less likely to seek medical help for non-mental health reasons than non-depressed individuals (even in case of severe diseases such as advanced cancer<sup>50</sup>), this probably explains the low prevalence of depression in this sample. Moreover, due to the cross-sectional nature of this study, we could not determine causality but only associations. Consequently, we could not ascertain whether skin diseases cause subjective well-being to decrease or whether individuals with low subjective well-being are more prone to developing skin diseases, e.g. due to psychoneuroimmunologic processes.<sup>51</sup> Finally, further variables influencing happiness, such as personality traits, which could potentially moderate the association between happiness and skin diseases,<sup>31</sup> had not been taken into account in this analysis. More detailed studies are needed to achieve a better understanding of their potential interaction with happiness in dermatologic patients.

#### Conclusion

The results of this study show that dermatologic patients tend to experience lower levels of happiness and especially PA compared to a healthy control group. As PA has been related to several desirable health outcomes, targeting PA in addition to the treatment of somatic symptoms is a promising approach for a holistic treatment of skin diseases.

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### Supporting information

Additional Supporting Information may be found in the online version of this article:

**Table S1.** Descriptive statistics for positive and negative affect using an alternative measuring tool (SPANES).

## 4. Paper II

### **Adding happiness to complement the Dermatology Quality of Life Index in Psoriasis and Atopic Dermatitis Healthcare**

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1

1 **Title:** **Adding happiness to complement the Dermatology Quality**  
2 **of Life Index in psoriasis and atopic dermatitis healthcare: a**  
3 **cross-sectional study**

4  
5 **Short title:** Adding happiness to complement the DLQI

6  
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30 **Figures:** 3

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32 **ABSTRACT**

33 *Background:* The Dermatology Quality of Life Index (DLQI) is the most commonly used quality  
34 of life questionnaire in dermatology.

35 *Objectives:* The aim of this study was to compare the DLQI to measures of well-being and  
36 general quality of life in patients with chronic inflammatory skin diseases.

37 *Material & Methods:* A cross-sectional study among patients with psoriasis and atopic  
38 dermatitis was conducted at a hospital in Munich, Germany (12/2017-04/2019). Participants  
39 filled in validated scales measuring happiness, quality of life, and DLQI.

40 *Results:* In 102 patients with chronic inflammatory skin diseases, the DLQI was associated only  
41 with physical quality of life and negative emotions but not psychological quality of life nor  
42 positive affect. The DLQI alone accounted for 26% of variance in general quality of life.  
43 Combining DLQI and happiness accounted for a total of 73% of variance, with both variables  
44 contributing to the model.

45 *Conclusion:* The DLQI alone only partially reflected well-being. Assessing happiness in addition  
46 to the widely used DLQI can contribute to a more comprehensive evaluation of well-being.

47

48 **Keywords:** atopic dermatitis, happiness, mental burden, psoriasis, quality of life, well-  
49 being

50

## 51 INTRODUCTION

52 Quality of life (QoL) is the most commonly used patient reported outcome in medicine (1).  
53 Since the World Health Organization (WHO) famously defined health “as a state of complete  
54 physical, mental, and social well-being and not merely the absence of disease” (2), the concept  
55 of QoL has received growing attention in medical research. From the 1970s to 1990s,  
56 numerous questionnaires measuring health-related QoL have been developed and  
57 introduced, including both generic and disease-specific tools (1, 3). In dermatology, the  
58 Dermatology Quality of Life Index (DLQI) (4) is the most well-established tool for measuring  
59 skin-related QoL and has proven its worth both in research and care (3).

60 Simultaneously to the growing efforts to measure well-being in medicine, a  
61 psychological sub-discipline researching well-being and human flourishing was formed in the  
62 second half of the 20<sup>th</sup> century and was given the name “Positive Psychology” in 1990 (5).  
63 Defining and measuring happiness is a major endeavor in positive psychology. Perhaps the  
64 most well-known conceptualization of happiness is Ed Diener’s “subjective well-being”, which  
65 describes “the cognitive and affective evaluation of life as a whole” (6, 7). According to this  
66 conceptualization, a person is happy if they experience a lot of positive emotions (positive  
67 affect, PA) while experiencing only few negative emotions (negative affect, NA) and if they are  
68 overall satisfied with their life (SWL) (7).

69 The first studies exploring happiness in dermatologic patients have shown that  
70 patients with skin diseases tend to be less happy compared to healthy controls and that  
71 patients with chronic inflammatory skin diseases, such as atopic dermatitis (AD) and psoriasis,  
72 are especially unhappy (8, 9), contributing to the considerable body of evidence  
73 demonstrating a large mental burden of these diseases (10-14). In an effort to meet the needs

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74 of these patients, the DLQI is commonly used in both clinical research and practice to reflect  
75 skin-related quality of life impairment (3). However, considering the WHO's holistic definition  
76 of health as a state of well-being raises the question whether examining only the DLQI, which  
77 was originally developed to measure disability (4), is sufficient in reflecting the patients'  
78 burden of disease or if other indicators of well-being should be added. To better understand  
79 the informative value of the DLQI for the evaluation of psychosocial needs, the aims of this  
80 study were to a) explore the association between the DLQI and happiness in patients with  
81 chronic inflammatory skin diseases and b) to put the results into perspective by making  
82 comparisons to a generic QoL assessment tool recommended by the WHO, the WHOQOL-  
83 BREF (15), and a control group.

#### 84 **MATERIALS AND METHODS**

85 Data were collected from December 2017 to April 2019. In- and out-patients with psoriasis  
86 and atopic dermatitis (AD) at the Department of Dermatology and Allergy of the Technical  
87 University of Munich, Germany, filled in a paper-based questionnaire (see supplementary  
88 material). All in- and outpatients aged 18 years or older who had been diagnosed with  
89 psoriasis or AD were eligible to participate. Exclusion criteria were the inability to provide  
90 consent and the inability to fill in the German questionnaire. A group of healthy controls (aged  
91 18 years or older) living in the greater Munich area, recruited with the help of a recruitment  
92 service (TestingTime, Zurich, Switzerland), filled in the same questionnaire. The study was  
93 conducted in accordance with the Declaration of Helsinki and was approved by the local ethics  
94 committee of the Technical University of Munich (reference number 424/17S).

95 *Questionnaire*

96 The questionnaire consisted of validated scales measuring happiness and QoL (see  
97 Supplement 1). The attending physician documented diagnoses, current treatment, and  
98 disease severity using the Psoriasis Area and Severity Index (PASI) for psoriasis and Scoring  
99 Atopic Dermatitis (SCORAD) (16) for AD. Disease severity was classified according to the  
100 respective manuals (mild: PASI/SCORAD < 10/25, moderate to severe: PASI/SCORAD ≥ 10/25)  
101 (17, 18). Current treatment was categorized as either systemic (e.g. immunomodulators, oral  
102 antihistamines) or non-systemic (e.g. topical steroids, phototherapy).

103 *Happiness* was operationalized as the three facets of subjective well-being(6): positive  
104 affect (PA), negative affect (NA), and satisfaction with life (SWL). The German version of the  
105 Scale of Positive and Negative Experience (SPANE) was used to measure PA and NA (19, 20)  
106 and consists of 12 items (6 representing PA, e.g. "pleasant", and 6 representing NA, e.g.  
107 "unpleasant"), each rated on a 5-point Likert scale ranging from 1 ("very rarely or never") to  
108 5 ("very often or always"). The two distinct subscales measuring PA and NA showed good  
109 reliability with Cronbach's alphas of .94 and .87, respectively. The items of each subscale were  
110 averaged to form an index if at least 80% of items were answered validly. SWL was measured  
111 using the German version of the Satisfaction With Life Scale (SWLS), a validated and widely  
112 used score consisting of 5 items (e.g. "in most ways my life is close to my ideals"), each  
113 measured on a scale from 1 ("strongly disagree") to 7 ("strongly agree") (21, 22). The SWLS  
114 showed good reliability in this sample with a Cronbach's alpha of .89. As with the SPANE scales,  
115 the items of the SWLS were averaged to form an index if at least 80% of items were answered  
116 validly.

117 *Quality of Life* was measured using the German version of the World Health  
118 Organization Quality of Life (WHOQOL)-BREF questionnaire (15, 23). The WHOQOL-BREF is a



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142 gender between the patients and the controls were analyzed using Student's t-test and chi-  
143 squared test. As the groups differed in terms of age, age was initially controlled for in all  
144 analyses. However, as age did not show to influence the results, it was omitted as a covariate  
145 to keep the models as simple as possible.

146 Descriptive statistics of happiness in both patient groups (psoriasis and AD) have been  
147 reported elsewhere in more detail (9). Student's t-tests were conducted to compare both  
148 patient groups in terms of happiness, DLQI, and QoL. As the analyses did not reveal major  
149 differences between the patient groups, all further analyses were conducted on the whole  
150 patient sample without further differentiation between the patient groups. Student's t-tests  
151 were used to compare patients and controls as well as compare patients receiving systemic  
152 treatment with patients receiving non-systemic treatment in terms of happiness and QoL.  
153 According to Diener's definition of subjective well-being (6) and following the methodology of  
154 existing research (24), a composite score of happiness was calculated as the mean of the z-  
155 standardized SWLS score, PA component of the SPANE, and inversely coded NA component of  
156 the SPANE. For the WHOQOL-BREF, the value domains were transformed to a scale of 0-100  
157 according to the manual (23) and prior to averaging them into an overall QoL index.

158 Linear regression models were used to explore the associations between happiness,  
159 QoL, and DLQI. Overall happiness, overall QoL, and the DLQI were explored as dependent  
160 outcomes. As independent variables, both the overall indices and the separate  
161 facets/domains were entered (in separate models) to allow a both detailed and concise  
162 analysis of the examined associations. Standardized regression coefficients ( $\beta$ ) are reported.

163 The level of significance was set at  $\alpha=.05$  for all analyses. All statistical analyses were  
164 conducted using IBM SPSS Statistics Version 26 (IBM Corporation, Armonk, NY, USA).

## 165 RESULTS

### 166 Study sample

167 In total, the data of 102 patients with chronic inflammatory skin diseases (52 with psoriasis,  
168 50 with AD) and 106 healthy controls were analyzed. The mean age of patients was  $47.4 \pm$   
169  $19.1$  years (range 18-86 years). Women constituted 43.1% of all participants. While a third of  
170 patients were only mildly affected, disease severity was moderate to severe in 67% of patients  
171 (psoriasis: 60.4%, AD: 73.3%;  $p=.187$ ). Almost half of all patients received a systemic treatment  
172 for their skin condition (46.4%, 4 cases with missing information). Patients with psoriasis and  
173 AD did not differ in terms of age or gender distribution ( $p=.493$  and  $p=.331$ , respectively).  
174 Participants of the control group were significantly younger than patients ( $m=38.4$ ,  $SD=13.4$   
175  $p<.001$ ) but showed a comparable gender distribution (49.1% women,  $p=.392$ ).

### 176 DLQI, QoL, and happiness

177 Patients with psoriasis and AD did not significantly differ in terms of DLQI ( $m=11.7$  vs.  
178  $m=14.4$ ,  $p=0.12$ ) nor happiness (PA:  $m=3.5$  vs.  $m=3.3$ ,  $p=.14$ ; NA:  $m=2.3$  vs.  $m=2.6$ ,  $p=.17$ ; SWL:  
179  $m=5$  vs.  $m=4.9$ ,  $p=.62$ ). In terms of QoL, patients with psoriasis reported better physical QoL  
180 than patients with AD ( $m=68.7$  vs.  $60.8$ ,  $p=.03$ ), but no difference was observed for the other  
181 QoL domains (mental:  $m=67.1$  vs.  $62.3$ ,  $p=.15$ ; social:  $m=69$  vs.  $71.3$ ,  $p=.6$ ; environmental:  
182  $m=75.6$  vs.  $74$ ,  $p=.54$ ).

183 In the overall patient sample, the mean DLQI score was  $13 \pm 8.6$  (Table 1). Overall, 54  
184 patients (52.9%) reported a very large or an extremely large effect of the respective skin  
185 disease on their lives according to the DLQI. For 18 patients (18.6%), the effect was moderate,  
186 and for 25 patients (24.5%) the skin diseases only had a small effect or no effect at all on their  
187 lives.

188 In terms of QoL, the patients scored lower than the controls in the physical and the  
189 psychological domains of QoL. Patients' results for the social and environmental domains were  
190 comparable to those for the control group. In terms of happiness, patients with psoriasis and  
191 AD reported lower levels of PA compared to the healthy control group, but they did not differ  
192 from the healthy controls in terms of NA nor SWL (Table 1).

193 Patients receiving systemic treatment showed a lower DLQI compared to patients  
194 receiving non-systematic treatment ( $m=10.7$  vs.  $m=15$ ,  $p=.02$ ), but they did not significantly  
195 differ in terms of happiness nor QoL as measured with the WHOQOL-BREF.

#### 196 **The association between DLQI, QoL, and happiness**

197 When differentiating the four domains of QoL, only the psychological ( $\beta=0.70$ ,  $p<.001$ ) domain  
198 of QoL significantly predicted happiness in the patient group. In contrast, the psychological  
199 domain ( $\beta=0.56$ ,  $p<.001$ ) and the social domain ( $\beta=0.22$ ,  $p=.001$ ) predicted happiness in  
200 healthy individuals (Table 2).

201 Linear regression furthermore revealed a significant association between overall QoL and  
202 DLQI ( $\beta=-.52$ ,  $p<.001$ ; Figure 1). More precisely, the DLQI significantly correlated only with the  
203 physical ( $\beta=-.42$ ,  $p=.002$ ) domain and not the psychological, social, and environmental  
204 domains of QoL. The DLQI furthermore negatively predicted overall happiness ( $\beta=-.49$ ,  $p<.001$ ;  
205 Figure 1). More precisely, an association between the DLQI with NA ( $\beta=.39$ ,  $p<.001$ ) and not  
206 with PA ( $\beta=-0.20$ ,  $p=.173$ ) nor with SWL ( $\beta=0.03$ ,  $p=.793$ ) was observed.

207 Happiness significantly predicted overall QoL in the patient group ( $\beta=0.84$ ,  $p<.001$ ) and  
208 control group ( $\beta=0.86$ ,  $p<.001$ ; Table 2). All three facets of happiness significantly predicted  
209 overall QoL in patients and controls, with SWL having the highest effect in both groups  
210 (patient:  $\beta=0.47$ ,  $p<.001$ ; control:  $\beta=0.40$ ,  $p<.001$ , Table 2).

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211            Figures 2 and 3 summarize the associations between DLQI, overall QoL, and happiness  
212 in the patient group. The DLQI alone ( $\beta=-0.52$ ,  $p<.001$ ) accounted for 26% of variance in overall  
213 QoL. Adding happiness to the model significantly increased the model's goodness of fit  
214 ( $p<.001$ ), with both happiness ( $\beta=0.77$ ,  $p<.001$ ) and DLQI ( $\beta=-0.13$ ,  $p=.031$ ) predicting overall  
215 QoL, resulting in a total of 73% of variance explained by the model.

## 216 **DISCUSSION**

217            This study explored the association between happiness, quality of life, and DLQI in  
218 patients with chronic inflammatory skin diseases. We found that while the DLQI correlated  
219 with both happiness and general QoL, it only reflected the negative emotions component of  
220 happiness (NA) and the physical domain of QoL, which in turn did not predict overall  
221 happiness. More precisely, the results show that the DLQI primarily measures physical  
222 impairment associated with negative emotions, which is in line with its original purpose as a  
223 "disability index" (4). Consequently, the DLQI alone showed to only partially reflect well-being.  
224 This conclusion is supported by a recent qualitative study, which found that patients felt that  
225 the DLQI did not adequately reflect their emotional burden (25). However, the DLQI is still  
226 used as the sole indicator of quality of life and well-being in many dermatological studies (26-  
227 28).

228            Happiness, in turn, strongly correlated with general QoL, suggesting that it measures  
229 well-being in a more comprehensive manner than the DLQI. Additionally, previous studies  
230 found that patients with psoriasis and AD experience less PA than healthy individuals, and that  
231 there may be an association between PA and disease severity (8, 9). PA has been linked to  
232 several desirable health outcomes such as better immune responses, improved health  
233 behavior, and quicker wound healing (29-31), which underlines the importance of PA for not

234 only mental but also physical health. As PA is not reflected in the DLQI, an important aspect  
235 of the burden of psoriasis and AD might have been overlooked in most clinical studies which  
236 only included the DLQI and not PA. Nevertheless, the DLQI contained information about  
237 general QoL, which was not provided by measuring happiness alone. Combining the frequently  
238 used DLQI with two short scales that measure happiness could provide a comprehensive  
239 assessment of QoL and well-being without the need for additional time-consuming QoL  
240 assessments.

241 While comparing DLQI and happiness is a novel approach, several validation studies  
242 have explored the association between DLQI and other generic QoL life measures (32). In  
243 patients with psoriasis and AD, the DLQI showed weak to moderate correlations with most  
244 generic QoL scores (e.g. the Nottingham Health Profile) (32, 33), which correspond with the  
245 comparably weak correlation between DLQI and overall QoL in this study. When considering  
246 the different domains of QoL, Shikiar et al. showed that the DLQI moderately correlated with  
247 both the mental and the physical components of the Short Form 36 (34), while in this study  
248 the DLQI was only associated with the physical and not the mental domain of QoL. However,  
249 Shikiar et al. used Pearson's correlation coefficients to independently compare the  
250 correlations between the single QoL domains and the DLQI (34), whereas in this study  
251 associations were explored using linear regression, controlling for all domains of QoL. When  
252 controlling for physical QoL, the association between DLQI and mental QoL seemingly  
253 disappeared.

254 The results of this study are subject to several limitations. First, as this is a cross-  
255 sectional study, we can only assess associations and not causality between happiness and QoL.  
256 Second, only patients with psoriasis and AD were included in the study. It is possible that the

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257 associations between the DLQI and happiness vary for different patient groups. More studies  
258 with larger and more diverse samples of dermatologic patients are needed to verify this.  
259 Furthermore, recruitment took place at only one university hospital in Munich, Germany.  
260 While the aim was to include all eligible patients, this was not always feasible and is why the  
261 sample is not necessarily representative of the respective patient collectives. Third, the  
262 findings only apply to the specific scales used in this study, namely the SPANE (19), the SWLS  
263 (21), the WHOQOL-BREF (15), and the DLQI (4). Although these scores are validated and well-  
264 established, it is possible that results vary when using other tools. This is especially true for  
265 happiness, as there is an ongoing debate about its operationalization in psychological research  
266 (35).

267 In conclusion, the results of this study show that the DLQI primarily measures physical  
268 impairment in patients with skin diseases and that it is predominantly associated with  
269 negative emotions. Therefore, the DLQI alone provides only a limited evaluation of well-being.  
270 Happiness is a very subjective measure of well-being with high individual relevance. It can  
271 complement the information provided by the DLQI and thus contribute to a comprehensive  
272 evaluation of well-being consistent with the WHO's definition of health. Happiness should be  
273 considered as a new patient reported outcome both in future research and in dermatologic  
274 care, as it could help to better understand the burden of skin diseases and guide therapy  
275 development and decisions. The development of a score assessing happiness in a  
276 comprehensive but uncomplicated manner would facilitate its application in clinical practice.

277

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373 **Figure Legends**

374

375 Figure 1: Associations between the Dermatology Quality of Life Index (DLQI) and a) happiness and b)  
376 World Health Organization Quality of Life (WHOQOL)-BREF.

377 Figure 2: Correlations between happiness, the Dermatology Quality of Life Index (DLQI) and World  
378 Health Organization Quality of Life (WHOQOL)-BREF in patients with psoriasis and atopic dermatitis.  
379  $R^2$  indicates the percentage of explained variance in the respective models (corrected  $R^2$ ).

380 Figure 3: Schematic illustration of the associations between happiness, quality of life (WHOQOL-BREF)  
381 and the Dermatology Quality of Life Index (DLQI). Happiness and QoL correlated more strongly than  
382 the DLQI and happiness/QoL, respectively. All three happiness facets correlated with QoL, with the  
383 strongest association observed for Satisfaction with Life (SWL). For QoL, only the psychological domain  
384 was associated with happiness. The DLQI correlated only with NA and physical QoL and thus only  
385 reflected a distinct part of well-being.

386

387 **Table Legends**

388

389 Table 1: Quality of Life and happiness in patients with psoriasis and atopic dermatitis compared to  
390 healthy controls and skin-related Quality of Life impairment (DLQI) in the patient group.

391 Table 2: Associations between World Health Organization Quality of Life (WHOQOL)-BREF, the  
392 Dermatology Quality of Life Index (DLQI), and happiness: results of linear regression models in  
393 patients with psoriasis and atopic dermatitis and healthy controls.

394 **Tables**395 **Table 1: Quality of Life and Happiness in patients with psoriasis and atopic dermatitis compared to healthy**  
396 **controls and skin-related Quality of Life impairment (DLQI) in the patient group**

	Patients N=102		Controls N=106		p <sup>a</sup>
	m / %	SD	m	SD	
Age	47.4	19.1	38.4	13.4	<.001
Gender (women)	43.1%	-	49.1%	-	3.92
Disease severity					
- mild	33.3%	-	-	-	-
- moderate to severe	66.7%	-	-	-	-
Systemic treatment	46.4%	-	-	-	-
DLQI <sup>b</sup>	13	8.6	-	-	
QoL <sup>c</sup> : physical	64.8	18.7	77.8	16.2	<.001
QoL <sup>c</sup> : psychological	64.7	16.9	68.9	15.9	.004
QoL <sup>c</sup> : social	70.1	21.1	68.6	22	.482
QoL <sup>c</sup> : environmental	74.8	13.7	74.6	15.8	.932
Overall QoL <sup>c</sup>	68.6	14.5	72.5	14.8	.057
Satisfaction with life <sup>d</sup>	5	1.2	4.8	1.3	.519
Positive affect <sup>e</sup>	3.4	0.9	3.8	0.8	.005
Negative affect <sup>e</sup>	2.4	0.9	2.3	0.8	.057
Overall happiness <sup>f</sup>	-0.1	0.9	0.1	0.9	.136

m=mean; SD=standard deviation; <sup>a</sup> Student's t-tests (for gender: Chi-Square test); <sup>b</sup> DLQI=Dermatology Quality of Life Index, possible range 0-30; QoL=Quality of Life, measured with the World Health Organization Quality of Life (WHOQOL)-BREF, possible range per domain: 0-10, overall QoL indicates average of all four domains. <sup>d</sup> measured using the Satisfaction with Life Scale, possible range 1-7; <sup>e</sup> measured using the Scale of Positive and Negative Experience, possible range 1-5; <sup>f</sup> satisfaction with life, positive and negative affect variables were z-standardized and averaged into an index, range in this sample: -3.03–1.47.

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**Table 2: Associations between World Health Organization Quality of Life (WHOQOL)-BREF, the Dermatology Quality of Life Index (DLQI), and happiness: results of linear regression models in patients with chronic inflammatory skin diseases and healthy controls**

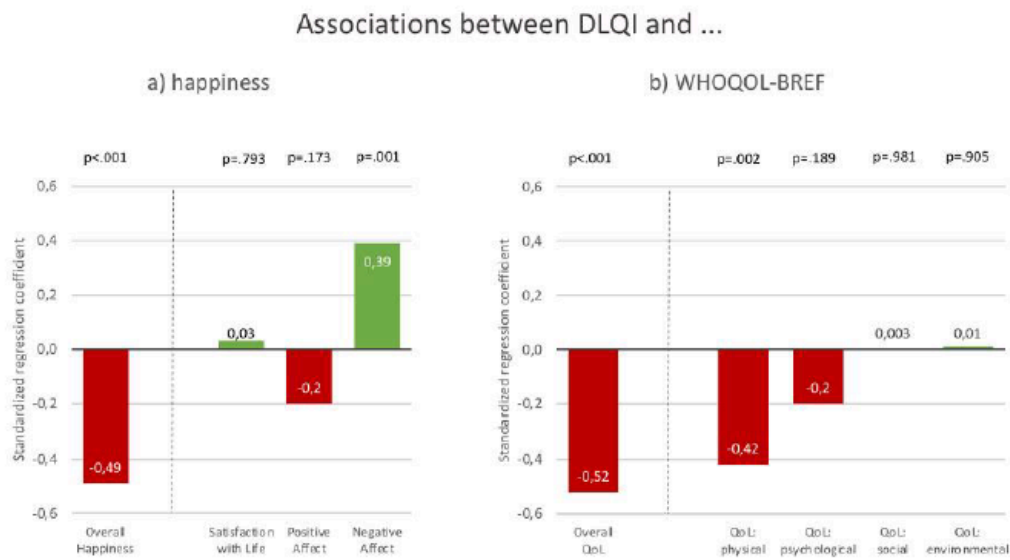
Predictors Model Summary	Patients N=102		Controls N=106	
DV: Overall Happiness				
QoL: physical ( $\beta$ , p)	0.08	.283	0.09	.224
QoL: psychological ( $\beta$ , p)	0.70	<.001	0.56	<.001
QoL: social ( $\beta$ , p)	0.11	.066	0.22	.001
QoL: environmental ( $\beta$ , p)	0.07	.250	0.13	.060
Model Summary (cR <sup>2</sup> )	0.77	<.001	0.77	<.001
DV: Overall QoL				
Overall happiness ( $\beta$ , p)	0.84	<.001	0.86	<.001
Model summary (cR <sup>2</sup> )	0.70	<.001	0.74	<.001
Satisfaction with life ( $\beta$ , p)	0.47	<.001	0.40	<.001
Positive affect ( $\beta$ , p)	0.27	.002	0.33	<.001
Negative affect ( $\beta$ , p)	-0.26	.001	-0.22	.006
Model summary (cR <sup>2</sup> )	0.71	<.001	0.74	<.001
DLQI ( $\beta$ , p)	-0.52	<.001	-	-
Model summary (cR <sup>2</sup> )	0.26	<.001	-	-
DLQI ( $\beta$ , p)	-0.13	.031	-	-
Overall happiness ( $\beta$ , p)	0.77	<.001	-	-
Model summary (cR <sup>2</sup> )	0.73	<.001	-	-

DV: dependent variable,  $\beta$ : standardized correlation coefficients, with positive and negative values indicating positive and negative associations, respectively, and higher absolute values indicating stronger associations between the DV and the predictor variables, p: p-value, QoL: Quality of life, cR<sup>2</sup>: corrected R<sup>2</sup>, indicating the corrected percentage of variance in the DV explained by the model.

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423 **Figures:**

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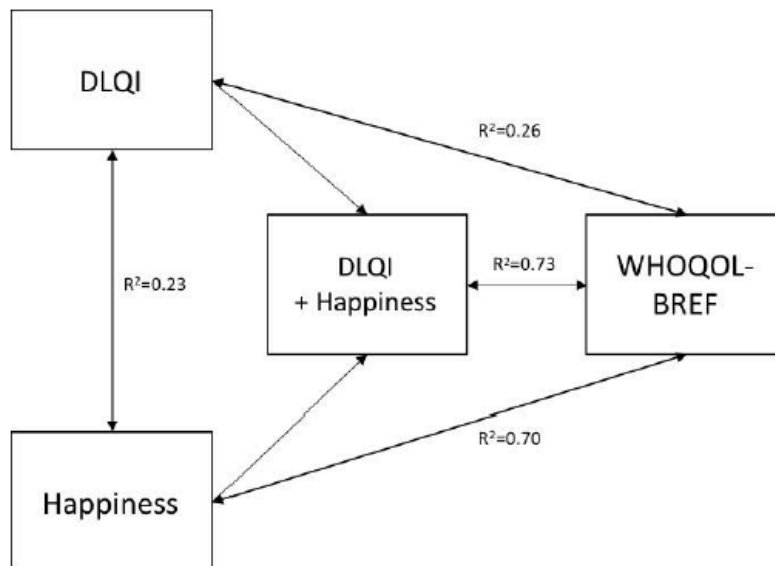


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426 Figure 1: Associations between Dermatology Quality of Life Index (DLQI) and a) happiness and b) World  
 427 Health Organization Quality of Life (WHOQOL)-BREF.

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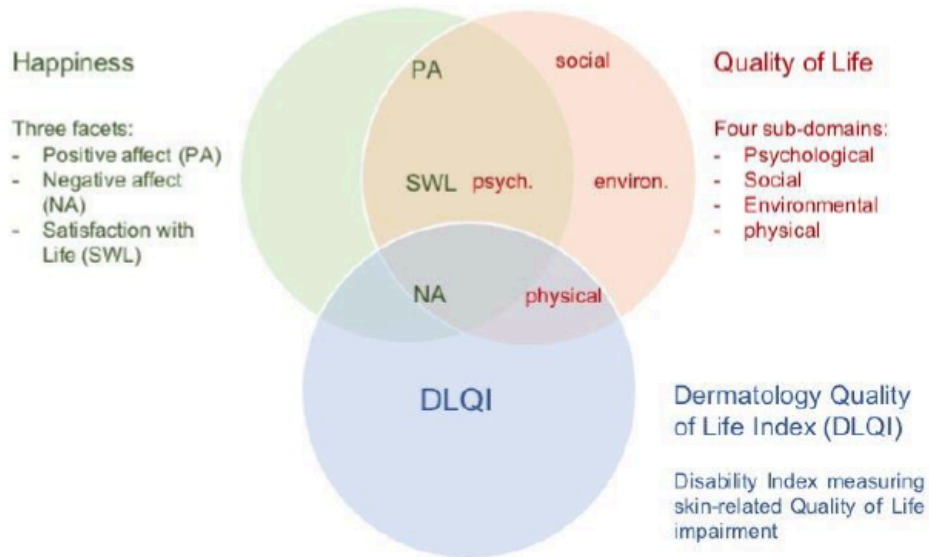
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431 Figure 2: Correlations between happiness, the Dermatology Quality of Life Index (DLQI) and World  
432 Health Organization Quality of Life (WHOQOL)-BREF in patients with psoriasis and atopic dermatitis.  
433  $R^2$  indicates the percentage of explained variance in the respective models (corrected  $R^2$ ).

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435

436 Figure 3: Schematic illustration of the associations between happiness, quality of life (WHOQOL-BREF)  
 437 and the Dermatology Quality of Life Index (DLQI). Happiness and QoL correlated more strongly than  
 438 DLQI and happiness/QoL, respectively. All three happiness facets correlated with QoL, with the  
 439 strongest association observed for Satisfaction with Life (SWL). For QoL, only the psychological domain  
 440 was associated with happiness. The DLQI correlated only with NA and physical QoL and thus only  
 441 reflected a distinct part of well-being.

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## 5. Paper III

### **Happiness and depression in psoriasis: a cross-sectional study in Germany.**

Schuster B, Peifer C, Ziehfrend S, Tizek L, Biedermann T, Zink A, Schielein MC

#### Quality of Life Research

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## Happiness and depression in psoriasis: a cross-sectional study in Germany

Barbara Schuster<sup>1,2</sup> · Corinna Peifer<sup>3</sup> · Stefanie Ziehfrend<sup>1</sup> · Linda Tizek<sup>1,2</sup> · Tilo Biedermann<sup>1</sup> · Alexander Zink<sup>1,4</sup> · Maximilian C. Schielein<sup>1,2,4</sup>

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### Abstract

**Purpose** Prior research on the psychological consequences of skin diseases has focused on assessing mental comorbidities. The aim of this study was to investigate subjective well-being in a large sample of individuals affected by psoriasis, a chronic inflammatory skin disease, and to explore the associations with depression and disease-related parameters such as disease severity.

**Methods** A cross-sectional online survey was conducted from March to June 2019. The link to the questionnaire was shared on websites and Facebook pages of psoriasis patient organizations and campaigns. Participants filled in validated scales measuring subjective well-being—operationalized as positive affect (PA), negative affect (NA) and satisfaction with life (SWL); and depression.

**Results** The data of 722 participants were analyzed. Exploratory factor analysis supported the differentiation of PA, NA, SWL, and depression as four different constructs. The respondents reported lower levels of PA than healthy individuals and judged themselves to be less happy and were less satisfied with their lives than the general population (except age group 65 + years). 40.3% of respondents were screened positive for depression. More severe psoriasis was associated with lower affective well-being and a higher risk for depression.

**Conclusion** The results of this study empirically supported the differentiation of subjective well-being and depression as different constructs in individuals with psoriasis, and underline the large mental burden of the disease which goes beyond a higher risk for depression. Measures of well-being should thus be incorporated in both research and clinical practice in patients with psoriasis in order to achieve a more comprehensive picture of the mental burden of this disease.

**Keywords** Psoriasis · Happiness · Subjective well-being · Depression · Mental health · Dermatology

### Plain English summary

Chronic skin diseases such as psoriasis can be a large mental burden for the affected individuals. Previous research has focused on assessing mental comorbidities such as depression in order to determine mental health in patients with skin diseases. However, according to the World Health Organization, there is more to being well than just not being ill. Thus, in order to fully capture the mental burden of skin diseases, both mental comorbidities and well-being need to be considered. Therefore, the aim of this study was to assess the mental burden of patients with psoriasis by not only measuring depression, but also happiness. We found that patients with psoriasis are less happy and have a higher risk for depression than healthy individuals. Treating skin symptoms can help the patients to be happier and less depressed.

Alexander Zink and Maximilian C. Schielein have contributed equally to this work.

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However, additional psychological care should be provided to patients with severe psoriasis in order to improve the patients' well-being.

## Introduction

Psoriasis is a common chronic inflammatory skin disease with an estimated prevalence of 1.2–3% in Germany [1, 2]. It is characterized by sharply demarcated red patches covered in typical whitish-silver scales, typically, but not only, located on knees and elbows as well as the scalp [3]. A large number of patients suffer from severe itch, and some of them, even pain [4, 5]. In addition, psoriasis is associated with a large number of comorbidities, ranging from arthritis and metabolic syndrome to depression [3, 6, 7]. As psoriatic lesions are often considered disfiguring by their social environments, many affected patients experience body image issues [8] and suffer from stigmatization and social exclusion [9, 10], resulting in a large psychological disease burden [7, 11, 12]. As a consequence, psoriasis has been recognized as a serious noncommunicable disease by the World Health Assembly in 2014 [13].

## Assessing the mental burden of psoriasis

Prior research on the psychological consequences of skin diseases has predominantly taken a pathogenic perspective, with most studies exploring mental comorbidities as markers of mental health. The results in general indicated a higher risk of depression, anxiety, and addiction for patients with psoriasis [6, 14–16]. However, in line with the World Health Organization (WHO) defining health as “a state of [...] well-being and not merely the absence of disease” [17], salutogenic approaches for measuring mental health in patients with skin diseases are necessary in order to achieve a holistic understanding of the mental burden [12, 18, 19]. Accordingly, over the last years, studies exploring well-being in individuals with psoriasis have emerged [20–24]. In general, most studies comparing individuals with psoriasis to healthy controls found that well-being was impaired in this patient group, underlining the large mental burden of psoriasis [20, 21, 25]. However, there is increasing evidence pointing toward a more complex relationship between psoriasis and well-being, which seems to be mediated by the subjective perception of the symptoms rather than the objective course of the disease [21, 26, 27].

## Subjective well-being and depression in psoriasis

Recently, happiness has been discussed as a patient reported outcome with high subjective relevance for patients with chronic skin diseases [20]. Although different definitions

exist [28], happiness is often conceptualized as subjective well-being [29]. According to Diener et al. [25], subjective well-being comprises an individual's “cognitive and affective evaluation of life as a whole”. Following this approach, subjective well-being is high if one experiences frequent positive affect (PA), infrequent negative affect (NA) and a general satisfaction with life (SWL) [25, 30]. So far, only a few studies exploring subjective well-being in patients with psoriasis exist [23, 24, 31, 32], with only one study comprehensively assessing all three facets of subjective well-being in a comparably small group of 52 patients with psoriasis [20]. The results of the existing studies suggest that psoriasis patients are impaired in terms of PA but not SWL compared to healthy individuals [20, 23, 24]. For NA, results are inconsistent [20, 23, 31].

Depression, in contrast, is a well-researched and generally accepted comorbidity of psoriasis which has been studied in several systematic reviews and meta-analyses [33, 34]. As depression is characterized by the reduced capability to experience positive emotions [35], the finding of low PA in patients with psoriasis could be explained by the higher prevalence of depressive symptoms in this patient group, which raises the question whether exploring affective well-being independently of depression does in fact provide new insights on the mental burden of psoriasis. Studies exploring both subjective well-being and depression in individuals with psoriasis are needed in order to be able to distinguish these two constructs in this patient group.

## Aim of the study

The aim of this study was thus to assess both depression and subjective well-being (operationalized as PA, NA, and SWL) in a large and diverse sample of individuals affected by psoriasis, and to explore associations between depression, subjective well-being, and patient characteristics in this group.

## Materials and methods

### Study population and recruitment

For this cross-sectional study, an online survey among individuals affected by psoriasis in Germany was conducted from March to June 2019. The link to the online questionnaire was shared on Germany's largest patients' organization webpage for psoriasis (“Psoriasis Netz”, [www.psoriasis-netz.de](http://www.psoriasis-netz.de)) and in the corresponding email newsletter and forum. In addition, the link was shared on other psoriasis-related Facebook pages (“Farbenhaut”, Technical University of Munich's University Hospital and the de-stigmatization campaign “Bitte berühren” by the Professional Association of the German Dermatologists). Only individuals affected

by psoriasis were eligible to participate, which was clearly stated on the first page of the online questionnaire and which had to be confirmed by the participants in the first question before they were able to continue with the rest of the questionnaire. Informed consent was obtained from all participants prior to inclusion. This study was conducted in accordance with the Declaration of Helsinki and was reviewed and approved by the local ethics committee of Technical University of Munich (reference 25/19 S).

### Study variables

The study questionnaire was developed by a multidisciplinary team including a dermatologist, a psychologist and two epidemiologists. It consisted of validated scales measuring subjective well-being and depression and additional questions on happiness in general and the evaluated impact of psoriasis. The survey was pre-tested by three psoriasis patients and, based on the resulting feedback, slightly modified. For the scales used in this study, internal consistency was considered “excellent” for Cronbach’s alpha values  $> 0.9$  and “good” for Cronbach’s alpha values  $> 0.8$  [36].

### Subjective well-being

Following Diener et al. [25], subjective well-being was operationalized as *PA*, *NA* and *SWL*. *PA* and *NA* were measured using the validated German version of the Scale of Positive and Negative Experience (SPANE) [37, 38]. We chose the SPANE over the more frequently used Positive and Negative Affect Schedule (PANAS) [39], as it assesses a wider range of positive and negative emotions than the PANAS, which focusses on high arousal emotions, and as it is coherent with Diener and colleagues conceptualization of subjective well-being [37]. The SPANE consists of two subscales measuring *PA* and *NA*, respectively. Each subscale consists of six adjectives, e.g., “pleasant” and “positive” for *PA* and “unpleasant” and “negative” for *NA*. The respondents were asked to indicate how often they had felt the respective feelings over the past two weeks on a 5-point scale from 1 (“very rarely or never”) to 5 (“very often or always”). The two distinct subscales, showed excellent and good reliability in this study with Cronbach’s alphas of 0.93 and 0.86 for *PA* and *NA*, respectively. For each subscale, the items were averaged to form an index. *SWL* was measured using the German version of the Satisfaction With Life Scale (SWLS) [40, 41]. The scale consists of five items (e.g., “in most ways my life is close to my ideal”), each rated on a 7-point scale from 1 (“strongly disagree”) to 7 (“strongly agree”). The scale showed excellent reliability in this study with a Cronbach’s alpha of 0.91 and the items were averaged to form an index. As some researchers argue that happiness is not in fact a multidimensional but a unidimensional construct [29], we

decided to also include a *heuristic measure of happiness*, meaning a single question asking participants for their overall happiness [20]. In order to achieve comparability with the general population, a single question from the European Social Survey was used: “Taking all things together, how happy would you say you are?” [42]. Respondents could give their answer on a 11-point scale from 0 (“extremely unhappy”) to 10 (“extremely happy”).

As an additional question on happiness in the context of psoriasis, the participants were asked for a *subjective evaluation of the impact of psoriasis on their own happiness* (“Do you think that your psoriasis has a negative impact on how happy you are?”—“No”, “Yes, a little”, “Yes, moderately”, “Yes, very much”).

### Depression

The WHO-5 Well-Being Index [WHO-5, 43] is a validated screening questionnaire for *depression*. It consists of five statements (e.g., “My daily life has been filled with things that interest me”), which respondents rate on a 6-point scale from 0 (“at no time”) to 5 (“all of the time”). The items showed good reliability in this study with a Cronbach’s alpha of 0.87. Following the instructions, the values of the items were added, resulting in an overall score ranging from 0 to 25, with lower scores indicating a higher risk for depression. Using a cut-off of  $\leq 7$ , the WHO-5 has shown good properties for the screening of major depression with a sensitivity of 94% and specificity of 78% [44]. Consequently, a score of  $\leq 7$  was considered a positive screening result for depression in this study.

### Participants’ characteristics

As possible parameters associated with happiness, data on *age* (in years), *gender*, *years since first diagnosis*, and *current treatment status* (currently receiving treatment vs. not receiving treatment) were collected. Subjective *general disease severity* was measured as “mild”, “moderate”, “severe”. In addition, data on current disease severity were collected using the same given options (“mild”, “moderate”, “severe”). Based on general and current disease severity, new dichotomous variables indicating current phases of *relative improvement* or *relative deterioration* compared to general disease severity were derived.

### Analysis

Prior to the analysis, data were checked for completeness (at least 80% of questions on happiness answered) and plausibility (e.g., data were considered implausible if the participants indicated a higher number of years since first diagnosis than age). As only very few participants did not meet our criteria

for completeness and plausibility ( $n = 8$ ), these cases were excluded from all further analysis rather than using imputation in order to keep the analysis as simple as possible. The remaining data were analyzed descriptively. Pearson's correlations were calculated in order to explore associations between the examined variables. Correlations were considered moderate for  $r > 0.5$  and strong for  $r > 0.7$  [45]. To further differentiate the four constructs PA, NA, SWL, and depression, exploratory factor analysis using Promin Rotation, an oblique rotation method appropriate for correlated variables, was conducted after checking the respective requirements were fulfilled. As all variables were measured on at least 5-step scales (with equal distances between the answer options and numbers suggesting equal steps between the categories), variables were treated as quasi-metric and Pearson's correlations were used for factor analysis. Following Guadagnoli and Velicer [46], factor loadings of 0.4 and higher were considered stable, which is why all smaller coefficients were suppressed. Factor retention was determined using parallel analysis [47]. However, as the retention of three factors, which was the number of factors suggested by parallel analysis, did not result in stable factor loadings for all examined items, we additionally conducted the analysis retaining one more factor, which was in line with the theoretical assumption of four differing constructs of PA, NA, SWL, and depression.

Means of PA, NA, heuristic happiness, SWL and WHO-5 were compared to norm data [41, 48, 49] or, in case of PA and NA, to data of a validation study [38] as norm data were not available. For both SWL and WHO-5, norm data had been collected in representative samples of the German general population (with the assistance of a demographic consulting company (SWL: mean age  $48.9 \pm 18.3$  years, 52.2% women; WHO-5: mean age 48.3 years, SD not indicated for the overall sample, 52.7% women) [41, 48]. For heuristic happiness, norm data for the German general population were collected within the European Social Survey (mean age  $48.2 \pm 18.1$  years, 49.9% women) [49]. For PA and NA, the data of the German validation study among a total of 498 participants recruited in university lectures ( $n = 264$ ) and via mailing lists/social media ( $n = 234$ ) were used as reference data (mean age not indicated, 18.5% < 20 years, 47.8% 20–29 years, 16.3% 30–39 years, 7.7% 40–49 years, 5.3% 50–59 years, 4.5% > 59 years, 75.1% women) [38]. As for PA, NA, and heuristic happiness the datasets of the reference samples were accessible, the group comparisons for these variables were conducted using ANCOVA and planned contrasts while controlling for sex and age. Adjusted means ( $m^a$ ) are reported. For SWL and WHO-5, the datasets of the reference populations could not be retrieved, which is why analyses were conducted stratified for age and sex, using Student's *t*-tests. The age groups

for these analyses were chosen to match the age groups reported in the respective reference samples. As Student's *t*-tests and ANCOVA have been shown to provide robust results even when normality and equal variance assumptions are violated [50–52], the analyses were conducted without prior verification of these assumptions.

Parameters associated with subjective well-being and a positive screening for depression were identified using multiple linear regression models and binary logistic regression, respectively. In all regression models, age, gender, years since first diagnosis, general subjective disease severity, improvement or deterioration of skin condition and current treatment status were entered as independent variables. As a result, adjusted raw (B) and standardized regression coefficients ( $\beta$ ) and Odds Ratios (OR) as well as corresponding 95%-confidence intervals (CIs) and the percentage of variance explained by each model (adjusted  $R^2$ ) are reported. As depression is characterized by the inability to feel happy [35], we conducted additional sub-group analyses in participants who did not receive a positive screening result for depression in order to explore differential findings for affective well-being which are not explained by the presence of depression.

The level of significance was set at  $\alpha = 0.05$  for all analyses. All statistical analyses except exploratory factor analysis were conducted using IBM SPSS Statistics Version 24 (IBM Corporation, Armonk, NY, USA). Exploratory factor analysis was conducted using FACTOR software [53].

## Results

In total, 730 participants completed the survey, with most users having participated via "Psoriasis Netz" (95%). Of those, eight were excluded prior to analysis due to insufficient ( $n = 3$ ) or implausible data (e.g., higher number of years since first diagnosis than age;  $n = 5$ ). Consequently, the data of 722 participants were analyzed. The mean age was 45.8 years with a standard deviation (SD) of 13.4 years and a range of 12–85 years (Table 1). 62.7% of participants were women. Almost all participants (98.6%) reported having been diagnosed with psoriasis by a medical doctor, and only ten participants (1.4%) had been given the diagnosis by an alternative practitioner or had self-diagnosed psoriasis. On average, the participants had been living with psoriasis for 20.6 years (SD = 14.5 years, range < 1–68). The majority of participants (55.7%) judged their psoriasis to be generally moderate, 34.3% severe, and only 10% mild. When asking for current disease severity, 24% of participants indicated mild, 52.8% moderate, and 23.3% severe. Comparing self-reported general and current disease severity, 31.7% of participants were in a phase of relative improvement, and 14.4%

**Table 1** Baseline patients' characteristics

Patients' characteristics	<i>N</i> = 722
Age in years (mean ± SD, range)	48.8 ± 13.4, 12–85
Gender	
- Male	269 (37.3%)
- Female	453 (62.7%)
Diagnosis	
- By medical doctor	712 (98.6%)
- By alternative practitioner or self-diagnosis	10 (1.4%)
Years since first diagnosis (mean ± SD, range)	20.6 ± 14.5, 0–68
General disease severity	
- Mild	72 (10%)
- Moderate	402 (55.7%)
- Severe	248 (23.3%)
Current disease severity	
- Mild	173 (24%)
- Moderate	381 (52.8%)
- Severe	168 (23.3%)
Phase of relative improvement of skin condition	229 (31.7%)
Phase of relative deterioration of skin condition	104 (14.4%)
Current treatment <sup>a</sup>	
- Dermatologist	446 (61.8%)
- Rheumatologist	127 (17.6%)
- General practitioner	117 (16.2%)
- Alternative care	23 (3.2%)
- No current treatment	179 (24.8%)

<sup>a</sup>Multiple answers were possible

of relative deterioration. One out of four participants (24.8%) was not currently receiving medical treatment for psoriasis. Of those who were receiving treatment, the majority (82.1%) were treated by a dermatologist, 23.3% by a rheumatologist, 21.5% by a general physician, and 3.2% received alternative care (e.g., by an alternative practitioner).

## Subjective well-being and depression

The variables intercorrelated moderately to strongly, with the strongest correlation observed for PA and heuristic happiness ( $r = 0.73$ ,  $p < 0.001$ ; Table 2) and the weakest correlation observed for SWL and NA ( $r = -0.56$ ,  $p < 0.001$ ). 40.3% of participants had a positive screening result for depression (41.9% of women and 37.5% of men). Nine out of ten participants (90.3%) stated that psoriasis had a negative impact on how happy they were (28.8% a little, 29% moderately, and 32.9% very much), and only 9.3% of participants stated that their psoriasis did not affect their happiness at all (Fig. 1).

Using exploratory factor analysis and parallel analysis for factor retention, three factors were identified (Fig. 2a). The first factor corresponded to the PA items of the SPANE, the second corresponded to the items of the SWLS, and the third factor corresponded to the NA items of the SPANE, with one item of the WHO-5 loading on the same factor. The remaining items of the WHO-5 did not show stable factor loadings on any of the three factors. When adding one more factor to be extracted, all items of SPANE, SWLS, and WHO-5 loaded on four separate factors corresponding to the four constructs PA, NA, SWL and depression (Fig. 2b).

Descriptive statistics of PA, NA, SWL, heuristic happiness, and WHO-5 are displayed in Table 3.

## Comparison with reference populations

Controlling for age and sex, participants in this study reported significantly less PA than the German validation sample ( $m^a = 2.98$  vs. 3.10,  $p = 0.003$ ) [38], but did not differ from them in terms of NA ( $m^a = 2.87$  vs. 2.88,  $p = 0.767$ ; Table 3). Furthermore, controlling for age and sex, participants in this study judged themselves less happy than the general population ( $m^a = 6.34$  vs. 7.26,  $p < 0.0001$ ). Stratifying for age and sex, participants of this study over all age groups and both men and women scored significantly lower on the WHO-5 than the general population (all  $p < 0.001$ ).

**Table 2** Correlation matrix

	PA	NA	SWL	Heuristic happiness	WHO-5 well-being index
PA	1				
NA	-0.669***	1			
SWL	0.652***	-0.564***	1		
Heuristic happiness	0.728***	-0.588***	0.629***	1	
WHO-5 well-being index	0.687***	-0.651***	0.630***	0.591***	1

Pearson's correlation coefficients are displayed

Asterisks indicate significant correlations; \* $p < 0.05$ , \*\* $p < 0.01$ , \*\*\* $p < 0.001$

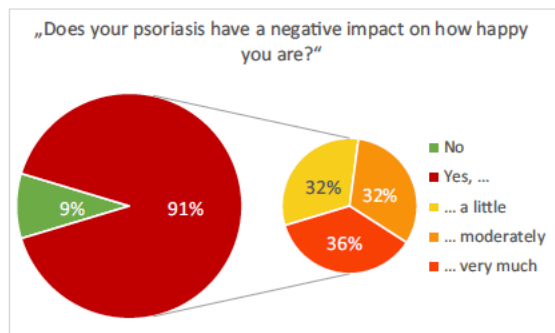


Fig. 1 Subjective evaluation of the impact of psoriasis on patients' happiness. *N*=721, one missing case

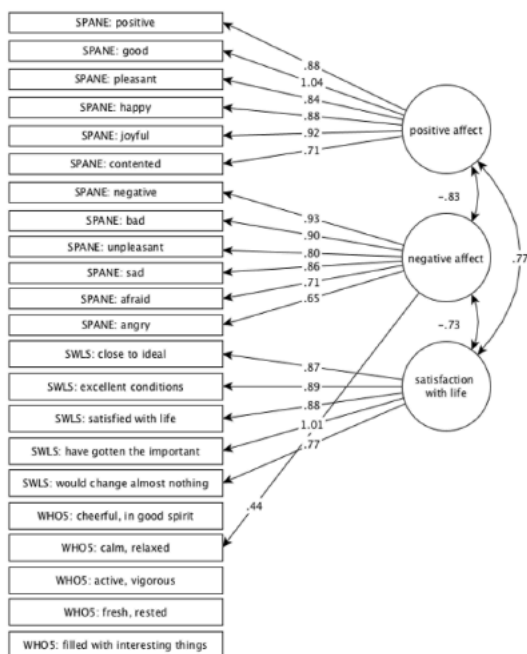
Furthermore, stratified analyses showed that participants under 65 years in this sample were less satisfied with their lives than the general population under 65 years (all  $p < 0.001$ ), while participants of 65 years and older did not differ from their peers in terms of SWL ( $p > 0.05$ ). Taking all age groups together, both men and women in this study reported lower SWL than men and women of the general population (both  $p < 0.0001$ ).

### Factors associated with subjective well-being and depression

#### PA

Higher age [ $B = -0.01$ , CI (-0.12; -0.002);  $\beta = -0.10$ , CI (-0.18; -0.03)], moderate [ $B = -0.33$ , CI (-0.55; -0.11);  $\beta = -0.18$ , CI (-0.31; -0.06)] and severe [ $B = -0.85$ , CI (-1.11; -0.59);  $\beta = -0.45$ , CI (-0.59; -0.31)] general disease severity as compared to mild disease severity and deterioration of skin condition [ $B = -0.32$ , CI (-0.53; -0.13);  $\beta = -0.13$ , CI (-0.2; -0.05)] were associated with lower levels of PA (Fig. 3). In contrast, a higher number of years since the first diagnosis [ $B = 0.11$ , CI (0.01; 0.02);  $\beta = 0.18$ , CI (0.10; 0.25)] and improvement of skin condition [ $B = 0.36$ , CI (0.21; 0.52);  $\beta = 0.06$ , CI (0.11; 0.27)] were associated with higher PA. The full model explained 10.7% of variance in PA ( $p < 0.001$ ). When including only patients without depression, the same parameters were associated with PA, but the model explained only 7.5% of variance ( $p < 0.001$ ).

(a) Factor retention based on parallel analysis



(b) Factor retention based on theoretical framework

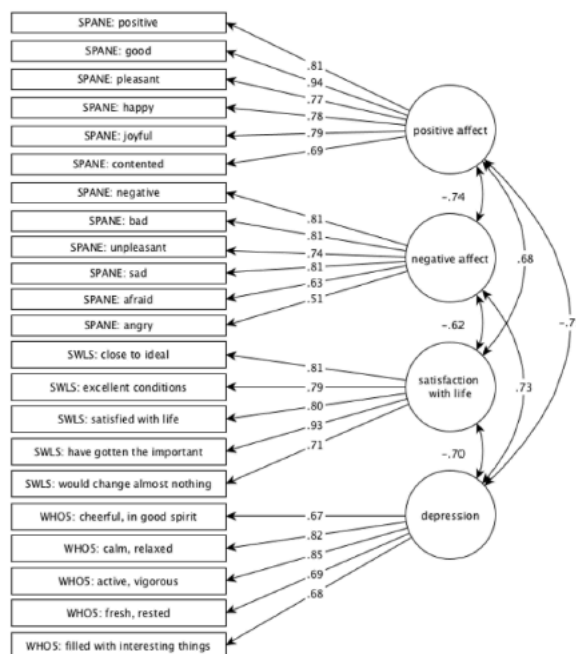


Fig. 2 Factor loadings of the examined variables of the SPANE, the SWLS, and the WHO-5 **a** on the three factors retained based on parallel analysis **b** on four factors corresponding to the four constructs

PA, NA, SWL, and depression. Arrows between factors show inter-factor correlations



## Quality of Life Research

**Table 3** Descriptive statistics of the examined variables in the study sample and the respective reference samples

	Mean $\pm$ SD	Adjusted mean/ means per group $\pm$ SD	Adjusted mean/means per group $\pm$ SD (reference)	Significance <sup>a</sup>
PA	2.96 $\pm$ 0.89	2.98	3.10	0.003
NA	2.83 $\pm$ 0.89	2.87	2.88	0.767
Heuristic happiness	5.32 $\pm$ 2.37	6.34	7.26	<0.001
SWL	4.27 $\pm$ 1.43			
- 14–24 years		19.63 <sup>b</sup> $\pm$ 6.24	24.77 <sup>c</sup> $\pm$ 6.11	<0.001
- 15–34 years		21.87 <sup>b</sup> $\pm$ 7.47	25.89 <sup>c</sup> $\pm$ 6.34	<0.001
- 35–44 years		20.77 <sup>b</sup> $\pm$ 7.66	25.01 <sup>c</sup> $\pm$ 6.37	<0.001
- 45–54 years		20.80 <sup>b</sup> $\pm$ 6.79	24.43 <sup>c</sup> $\pm$ 6.81	<0.001
- 55–64 years		21.28 <sup>b</sup> $\pm$ 7.28	24.15 <sup>c</sup> $\pm$ 6.32	<0.001
- 65–74 years		23.73 <sup>b</sup> $\pm$ 6.13	25.30 <sup>c</sup> $\pm$ 5.52	0.059
- > 74 years		24.56 <sup>b</sup> $\pm$ 3.78	25.06 <sup>c</sup> $\pm$ 5.93	0.802
- Men		20.86 <sup>b</sup> $\pm$ 7.40	25.12 $\pm$ 6.32	<0.001
- Women		21.62 <sup>b</sup> $\pm$ 7.05	24.67 $\pm$ 6.20	<0.001
WHO-5 well-being index	9.86 $\pm$ 5.27			
- < 41 years		9.73 $\pm$ 4.96	18.36 $\pm$ 4.80	<0.001
- 41–60 years		9.41 $\pm$ 5.30	17.49 $\pm$ 4.88	<0.001
- > 60 years		11.9 $\pm$ 5.65	16.70 $\pm$ 5.13	<0.001
- Men		10.34 $\pm$ 5.61	18.15 $\pm$ 4.90	<0.001
- Women		9.58 $\pm$ 5.05	17.07 $\pm$ 4.98	<0.001

<sup>a</sup>ANCOVA and Student's *t*-tests, respectively<sup>b</sup>For stratified analysis items were summed up (instead of averaged) in order to allow for comparison with reference data<sup>c</sup>Calculated based on weighted means for men and women

## NA

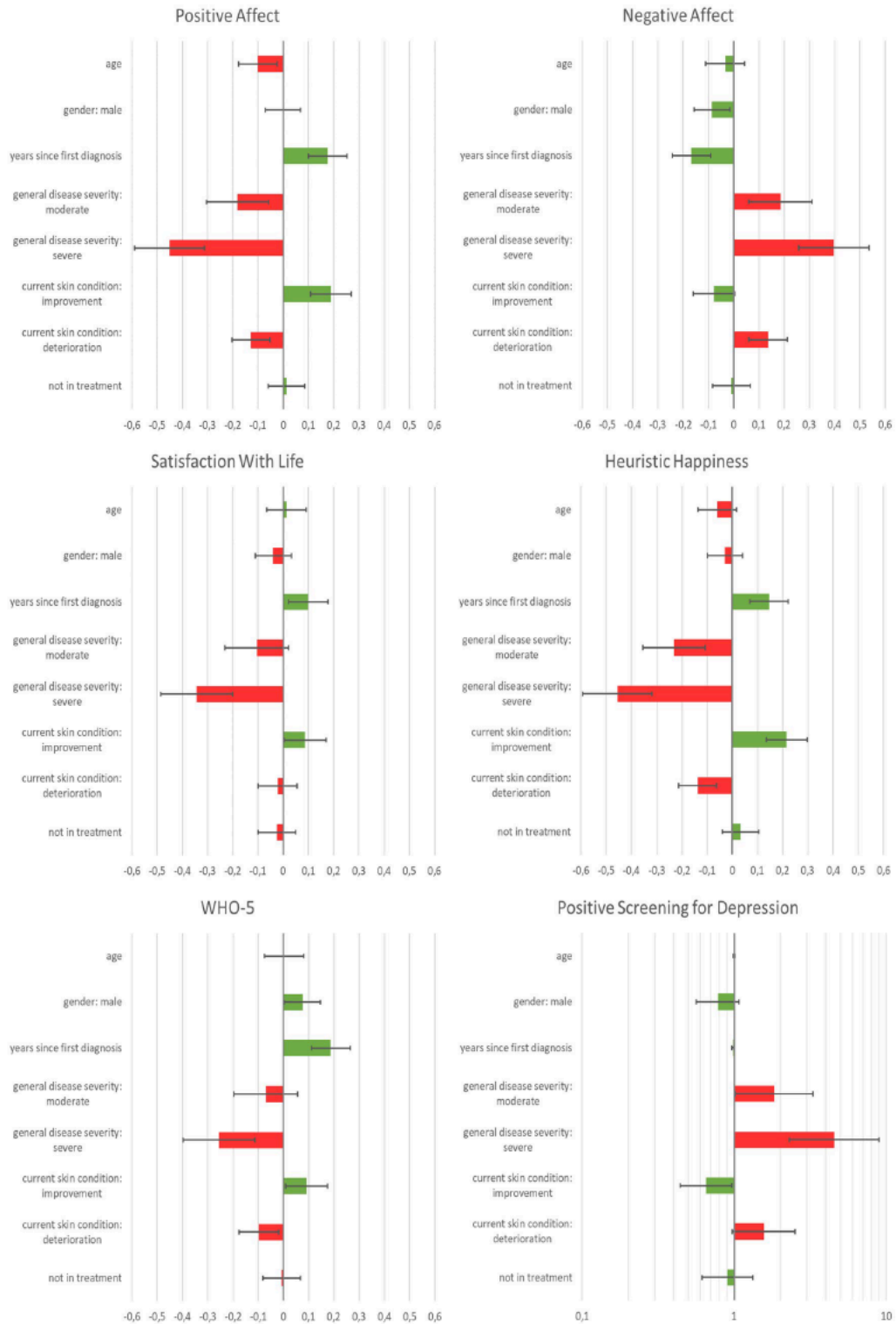
NA was higher in participants with moderate [B = 0.33, CI (0.11; 0.55);  $\beta$  = 0.19, CI (0.06; 0.31)] and severe [B = 0.74, CI (0.48; 1);  $\beta$  = 0.40, CI (0.26; 0.54)] general disease severity and those who were currently in a phase of deterioration of the skin condition [B = 0.34, CI (0.15; 0.54);  $\beta$  = 0.14, CI (0.06; 0.21); Fig. 3]. In contrast, being male [B = - 0.16, CI (- 0.29; - 0.03);  $\beta$  = - 0.09, CI (- 0.16; - 0.02)] and living with psoriasis for a longer time [B = - 0.01, CI (- 0.02; - 0.006);  $\beta$  = - 0.17, CI (- 0.24; - 0.09)] were associated with less NA. The full model explained 9.1% of variance in NA ( $p$  < 0.001). When including only patients without depression, the association remained significant for moderate and severe general disease severity and years since the diagnosis of psoriasis ( $p$  = 0.008;  $p$  = 0.001;  $p$  < 0.001, respectively), but the associations with gender and deterioration of the skin did not reach significance anymore ( $p$  = 0.071 and  $p$  = 0.085, respectively). Similar to PA, the model explained slightly less variance (7.6%,  $p$  < 0.001) in participants without depression compared to the whole sample.

## SWL

Severe general disease severity was associated with lower levels of SWL [B = - 1.04, CI (- 1.47; - 0.61);  $\beta$  = - 0.34, CI (- 0.49; - 0.20); Fig. 3]. More years passed since the first diagnosis [B = 0.01, CI (0.002; 0.02);  $\beta$  = 0.10, CI (0.02; 0.18)] and current improvement of skin condition [B = 0.27, CI (0.01; 0.52);  $\beta$  = 0.09, CI (0.00; 0.17)] were associated with higher SWL. The full model explained 5.5% of variance ( $p$  < 0.001) in SWL. When excluding participants with depression, only female gender was significantly associated with SWL ( $p$  = 0.049). Overall, the model did not show a good fit for this sub-sample (adjusted  $R^2$  = 0.013,  $p$  = 0.099).

## Heuristic happiness

Patients who were in general moderately [B = - 1.11, CI (- 0.17; - 0.51);  $\beta$  = - 0.24, CI (- 0.36; - 0.11)] or severely [B = - 2.28; CI (- 2.97; - 1.59);  $\beta$  = - 0.46, CI (- 0.60; - 0.32)] affected or who were currently in a phase of deterioration [B = 1.1, CI (0.69; 1.51);  $\beta$  = - 0.14, CI (- 0.22; - 0.07)] judged themselves to be less happy



**Fig. 3** Determinants of happiness and depression in 722 individuals with psoriasis. Adjusted standardized regression coefficients (for positive screening of depression: OR on log scale) with corresponding confidence intervals are given. Factors associated with higher well-being are displayed in green and those associated with lower well-being are displayed in red. Happiness was operationalized as subjective well-being, consisting of PA, NA, and SWL, and as a heuristic evaluation of happiness. A screening result for depression was considered positive if a participant scored  $\leq 7$  in the WHO-5. Asterisks indicate significant correlations. \* $p < 0.05$ , \*\* $p < 0.01$ , \*\*\* $p < 0.001$ . (Color figure online)

(Fig. 3). In contrast, those who had been living with psoriasis for more years [B = 0.02, CI (0.01; 0.04);  $\beta = 0.14$ , CI (0.07; 0.22)] or who were in a current phase of improvement of skin condition [B = 1.1, CI (0.69; 1.51);  $\beta = 0.22$ , CI (0.14; 0.30)] judged themselves to be happier. The full model explained 10.2% of variance in heuristic happiness. All parameters associated with heuristic happiness remained significant when excluding patients with depression, and the model explained 6.9% of variance in heuristic happiness ( $p < 0.001$ ).

### Depression

Participants with severe psoriasis [B = - 2.84, CI (- 4.41; - 1.27);  $\beta = - 0.26$ , CI (- 0.40; - 0.11)] and currently going through a phase of deterioration [B = - 1.46, CI (- 2.62; - 0.3);  $\beta = - 0.10$ , CI (- 0.18; - 0.02)] scored lower in the WHO-5 questionnaire, which indicates higher levels of depressive tendencies (Fig. 3). In contrast, men [B = 0.82, CI (0.04; 1.6);  $\beta = 0.08$ , CI (0.01; 0.15)], participants living with psoriasis for a longer time [B = 0.07, CI (0.04; 0.1);  $\beta = 0.19$ , CI (0.11; 0.27)] and those currently experiencing improvement of skin condition [B = 1.04, CI (0.11; 1.98);  $\beta = 0.09$ , CI (0.01; 0.18)] scored higher in the WHO-5. The full model explained 6.6% of variance in the WHO-5 score ( $p < 0.001$ ). Moderately affected participants had a 1.8-fold [CI (1.02; 3.32)] and severely affected participants even had a 4.6-fold [CI (2.31; 9.03)] chance of being screened positive for depression compared to mildly affected participants (Fig. 3). Living with psoriasis for a year longer [OR 0.98, CI (0.97; 0.99)] and current improvement of skin condition [OR 0.65, CI (0.44; 0.97)] were both associated with a decreased chance of a positive screening for depression. The full model explained 8.6% of variance in screening results ( $p < 0.001$ ).

### Discussion

This study examined both subjective well-being and depression in patients with psoriasis, following the WHO's demand for a holistic approach of measuring mental health in this patient group [12]. In a nationwide sample of 722 individuals

affected by psoriasis in Germany, subjective well-being and more precisely PA and SWL were low compared to the general population or healthy reference populations, and 40% of participants were screened positive for depression. The main risk factor for low subjective well-being and depression was general disease severity, but also deterioration of the skin was negatively associated with the affective component of subjective well-being, while phases of improvement of skin condition were associated with more PA and higher SWL. Additionally, the longer participants had been living with psoriasis, the higher was their subjective well-being and the less depressed they were.

### Differentiating depression and well-being

Due to the simultaneous assessment of depression and subjective well-being, this study allowed to examine the association between these constructs in individuals affected with psoriasis. As depression, among other symptoms, is characterized by the absence of PA [35], the constructs of depression and subjective well-being/positive affect are bound to overlap per definition, which is reflected by overlapping items of the respective measurement tools (especially question 1 of the WHO-5 and the PA items of the SPANE) [54]. However, depression is also characterized by other symptoms such as lack of energy and decreased activity which are not covered by PA/subjective well-being. Accordingly, exploratory factor analysis revealed that all items of SPANE, SWLS and WHO-5 loaded on four different factors corresponding to PA, NA, SWL and depression, supporting the methodological approach of differentiating depression and subjective well-being. This is especially interesting as the empirical differentiation of subjective well-being and depression, which has been successfully shown in some studies [55], has proven difficult in others [56]. The results of this study are in line with the postulate of the WHO that well-being is more than the absence of illbeing (see also Lukat et al. [57]). Thereby, they underline the WHO's demand to including measures of well-being in clinical research and practice in individuals with psoriasis in order to achieve a more comprehensive understanding of the mental burden of this disease [12].

### Subjective and objective disease severity

Subjective general disease severity was associated with all facets of subjective well-being in this study. In contrast, two previous studies have only found associations between disease severity and single facets of subjective well-being (NA: Martin-Brufau et al. [23]; PA: Schuster et al. [20], respectively). These inconsistencies in the findings might be owed to the different modalities of assessing disease severity employed in the different studies: Previous studies

have shown that the subjective perception of the symptoms might be more important for individual well-being than the objective course of the disease [21, 26, 27]. While the two mentioned studies used either physician assessed disease severity [20] or a validated self-assessment tool for disease severity [23], a very subjective approach was used in the present study, with participants rating their disease severity as “mild”, “moderate” or “severe”. This approach might explain why the associations between well-being and disease severity were especially pronounced in this study. More research is needed in order to shed light on the (different?) roles of subjective and objective disease severity for well-being and mental health. An alternative explanation could be that the small sample size of 52 patients in the study of Schuster et al. [20] prevented the detection of associations between disease severity and NA and SWL observed in this study.

### General and current disease severity

Interestingly, high general disease severity remained strongly associated with low subjective well-being and depression even when current skin condition was taken into account. While improvement of the skin was associated with higher well-being, the negative association between general severe disease severity and well-being was more pronounced. These findings implicate that by relieving skin symptoms, physicians might be able to help patients feel happier and, as reported before [15, 16], reduce the risk for depression. However, especially in more severe forms of psoriasis, well-being might remain low even if the symptoms improve. This supports the idea that documenting the PeakPASI, meaning the highest PASI ever recorded per patient, might provide additional information about the mental burden of psoriasis compared to just referring to the “snapshot” PASI at a single visit [58].

### The role of online recruitment

The participants in this sample scored lower than the German norm in SWL and reported less PA than a healthy reference sample. While the findings of impaired PA, but not increased NA, is in line with previous findings [20], the reported values for PA, NA and SWL were even more unfavorable than in a previous study among patients with psoriasis [20]. Also, we found a high rate of positive screening for depression of 40%. An explanation for these findings could be that the recruitment strategy might have led to a sample which was especially impaired in terms of well-being and depression. This would suggest that especially unhappy individuals turn to (online offers of) self-help organizations like the one used for recruitment in this study, which would underline the great importance these entities have for the

comprehensive treatment of individuals with psoriasis. In contrast, however, it is also possible that, due to the anonymity, the participants in this online survey were more honest and open about their well-being than in previous studies, which have mostly taken place in medical settings [20, 24, 32]. Thus, social desirability might have led to an underestimation of happiness and well-being in previous studies.

### Limitations

The results of this study are subject to several limitations. First, as the survey was conducted online, selection bias must be considered, which was already discussed in the above paragraph. Second, while all participants had to actively indicate that they were indeed affected by psoriasis prior to taking part in the survey, the online setting did not allow to confirm the diagnosis. Third, due to time limitations in the survey, which was designed to take only a few minutes to complete in order to achieve a high number of participants, disease severity was not assessed using validated scales or clinical scores but as “mild”, “moderate” and “severe”. Future studies should explore whether the findings of this study can be replicated when using validated scores for the assessment of subjective or even objective disease severity like PASI. Also, as this is a cross-sectional study, we could not determine causality but only association. Finally, future studies should explore the role of personality traits and other possible moderator variables such as marital status for the examined associations, as they have not been considered in this study.

### Strengths

This study followed a holistic approach of assessing mental health in individuals with psoriasis by measuring not only depression, but also subjective well-being in a large sample of affected individuals. Thus, the study does not only contribute to the growing body of research exploring well-being in psoriasis healthcare, it furthermore adds empirical evidence supporting the differentiation of depression and well-being as two related constructs. Furthermore, the study was designed in a way to not only include patients in the medical setting, but also affected individuals who were not currently part of the health care system, which is in line with the WHO’s demand for “people centered care” instead of “patient centered care” [59]. Thus, by looking at people with psoriasis in general instead of focusing on patients only (= people who are currently receiving medical care), the study provides a more comprehensive picture of mental health in individuals with psoriasis regardless of treatment status than prior studies which were mostly conducted in medical settings. Finally, the online setting of this study allowed the participants to fill in the questionnaire in an

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anonymous setting, which might help to reduce social desirability bias as compared to paper-based questionnaires at doctors' offices or clinics which are frequently employed in epidemiological research.

### Future research

Interventions targeting NA (especially stress) have been successfully conducted in patients with psoriasis [60, 61]. As this and previous studies have shown that PA seems to be especially impaired in patients with psoriasis [20], and as PA has been linked to several desirable health outcomes like better immune reactions, faster skin barrier recovery, and better cardiovascular health [62, 63], future research should evaluate whether interventions targeting PA could be also or maybe even more beneficial for improving the patients' well-being beyond the treatment of skin symptoms. Furthermore, future research should explore the mechanisms behind the presumably beneficial effect of skin treatment on well-being, as it could be moderated and potentially improved by additional factors such as time to treatment response [26].

### Conclusion

In this large sample of 722 patients with psoriasis, participants reported lower subjective well-being than the German general population or healthy reference populations. General disease severity was associated with both low subjective well-being and depression, even in phases of improvement of the skin condition. While improvement of skin condition, which can be achieved by an effective treatment of skin symptoms, was associated with higher well-being and less depression, the negative associations between severe general disease severity and well-being and depression, respectively, were even more pronounced. As subjective well-being and depression have been identified as two differential constructs in this study, measures of well-being should be increasingly incorporated in psoriasis health care.

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AZ: supervision; MCS: project administration. All authors have read and agreed to the published version of the manuscript.

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**Data availability** Data are available at <http://dx.doi.org/10.23668/psycharchives.4451>.

**Code availability** Code is available upon request.

### Declarations

**Conflict of interest** The authors declare that they have no conflict of interest.

**Ethical approval** This study was reviewed and approved by the local ethics committee of Technical University of Munich (reference 25/19 S).

**Consent to participate** All participants provided informed consent prior to participation.

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