#### RESEARCH



# Prevalence and severity of anxiety in cancer patients: results from a multi-center cohort study in Germany

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#### Abstract

**Purpose** Anxiety is an accompanying symptom in cancer patients that can have a negative impact on patients. The aim of the present analyses is to determine the prevalence of anxiety, taking into account sociodemographic and medical variables, and to determine the odds ratio for the occurrence of anxiety in cancer patients compared to general population.

**Methods** In this secondary analyses, we included 4,020 adult cancer patients during and after treatment from a multi-center epidemiological study from 5 regions in Germany in different treatment settings and a comparison group consisting of 10,000 people from the general population in Germany. Anxiety was measured with the Generalized Anxiety Disorder (GAD-7) questionnaire. In multivariate analyses adjusted for age and sex, we calculated the odds of being anxious.

Results The prevalence of anxiety was observed to be 13.8% (GAD- $7 \ge 10$ ). The level of anxiety was significant higher for patients in rehabilitation, compared to patients during inpatient and outpatient treatment (p = .013). Comparison with the general population yielded a 2.7-fold increased risk for anxiety among cancer patients (95% CI 2.4–3.1; p < .001). Patients with bladder cancer (OR, 5.3; 95% CI 3.0–9.4) and testicular cancer (OR, 5.0; 95% CI 2.1–12.1) showed the highest risk of having high levels of anxiety.

**Conclusion** The results highlight the importance of identifying anxiety in cancer patients.

Keywords Anxiety · Cancer · Oncology · Anxiety symptoms · GAD-7

#### Introduction

A critical life event may increase the risk for psychological distress and anxiety (Hassanzadeh et al. 2017). Therefore, anxiety is also a well-known accompanying phenomenon in cancer patients (Kapfhammer 2015; Bates et al. 2017; Pitman et al. 2018). A meta-analysis of studies from different countries found a prevalence of anxiety disorders in 10.3% (95 CI 5.1–17.0) of patients for hematology and oncology (Mitchell et al. 2011). This same meta-analysis demonstrated

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anxiety disorders in 9.8% (95% CI 6.8–13.2) of palliative patients. Another review reported both a pooled adjusted 4-week prevalence of anxiety disorders in cancer patients of 13.5% (95% CI 7.1–24.3) based on 10 studies and a lifetime prevalence of 30.5% (95% CI 28.0-33.0) based on 2 studies for Germany (Vehling et al. 2012). Both reviews integrated studies that assessed anxiety disorders by structured clinical interviews according to Diagnostic and Statistical Manual of Mental Disorders (DSM) or International Statistical Classification of Diseases and Health-Related Problems (ICD-10). It is known from other studies that results depend on the method chosen and vary accordingly (Krebber et al. 2014). In contrast, studies using validated questionnaires for self-assessment of anxiety showed different results. A study conducted in Utrecht observed anxiety symptoms in 22% of 2,144 inpatients (Van Den Brekel et al. 2020). In patients treated exclusively with symptomatic palliative care, the prevalence raised up to 36%. Predictors proved to be female gender, younger age, depressed mood, sleeping problems, and dyspnea. Maass et al. (2015) concluded from a review of studies in women with breast cancer that



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the prevalence of anxiety ranges from 17.9 to 33.3% and remains so in breast cancer survivors. Anxiety in patients with cancer in low- and lower-middle-income countries was addressed by authors in another review (Walker et al. 2021). They observed a prevalence between 5.6 and 89.2%. These were often associated with advanced disease and low levels of education. However, these results have to be considered cautiously against the background of different questionnaires and different thresholds.

Why is anxiety a non-negligible factor in the context of oncological diseases? Anxiety and a correspondingly elevated physiological level of arousal increase functions that can be influenced by the autonomic nervous system, such as pain, nausea and vomiting. This in turn can lead to a renewed increase in anxiety. Anxiety can lead to behavioral paralysis and patients are unable to apply adequate coping strategies. Therefore, anxiety can affect compliance with treatment (Housman et al. 2021). A correlation between anxiety and increased side effects during chemotherapy could be observed (Mahdizadeh et al. 2019). A study in cancer patients on oral targeted or chemotherapy reported an association between fatigue and anxiety (Poort et al. 2020). Furthermore, it could be shown that anxiety influence the functional outcome (Dinesh et al. 2021). A review of international studies revealed that clinically diagnosed anxiety disorders and anxiety defined by scales were related with higher cancer-specific mortality as well as poorer survival (Wang et al. 2020). A study in Germany examined 436 female patients with breast cancer concluded that increased anxiety was associated with greater dissatisfaction with information both at baseline and 12 months follow up (Faller et al. 2017). Another study from Germany showed a similar result. Here, in a multi-center longitudinal study, 1,398 patients with various cancer diagnoses were surveyed regarding their experience of anxiety at three different points in time (baseline, 6 months and 12 months follow up). As a result, it was shown that increased anxiety at all time points was associated with greater dissatisfaction with information perceived (Goerling et al. 2020). Analyses of the prevalence of anxiety in long-term cancer survivors yielded 17.9% (95% CI 12.8-23.6) (Mitchell et al. 2013). Out of 1154 survivors, 21% reported a score of  $\geq 8$  on the anxiety scale measured with the Hospital Anxiety and Depression Scale (HADS) 12 months after diagnosis (Boyes et al. 2013). Anxiety has a negative impact on quality of life after cancer (Saevarsdottir

Many previous studies have targeted patient populations with common tumor diseases such as breast or prostate cancer or specific stages of cancer treatment. We therefore aim to estimate the prevalence of anxiety based on a large multicenter epidemiological study with multiple tumor locations. We analyze differences between individual tumor entities during and after treatment and we compare the anxiety of

these patients with the age- and gender-adjusted general population.

#### **Methods**

Here, we present data from a secondary analysis of a multicenter epidemiological cross-sectional study where we investigated the prevalence of mental disorders, psychosocial distress and need for psychosocial support in cancer patients (Mehnert et al. 2014). The sample is composed of oncology patients receiving treatment in both inpatient and outpatient settings. We also included patients in cancer rehabilitation. Patients with malignant tumor across all entities and disease stages were included, and stratified by nationwide incidence of cancer diagnoses. The recruitment took place consecutively from acute care hospitals, outpatient cancer care facilities, and cancer rehabilitation centers in Germany. All patients provided written informed consent. The Ethics Committees of all participating centers obtained positive ethical approval. The study complied with the Declaration of Helsinki as well as the terms of data protection and privacy laws. The study protocol was described in detail elsewhere (Mehnert et al. 2012).

### **General population data**

To compare anxiety from cancer patients with the general population, we used data from a large population-based study, including 10,000 people in Germany (Loeffler et al. 2015; Hinz et al. 2017). This study contained an age- and gender-stratified random selection of inhabitants.

#### **Measures**

For the demographic variables age, sex, partnership, years of education, and work situation of cancer patients were documented. Medical data were obtained from medical records and included cancer diagnosis (ICD-10, WHO; 2004), tumor stage (UICC TNM classification; Sobin et al. 2009), metastases, cancer disease condition, treatment intention (curative/palliative), and completed treatment (surgery, radiation, chemotherapy, hormone treatment). Furthermore, we obtained the performance status (ECOG; Oken et al. 1982). The score can range from 0 (asymptomatic) to 4 (bedbound).

Anxiety was measured using the German version of GAD-7 (Spitzer et al. 2006; Löwe et al. 2008). This is a one-dimensional instrument created to detect symptoms of generalized anxiety disorders defined in the DSM-IV (APA 2000), and has excellent reliability, as well as criterion, construct, factorial, and procedural validity. Core symptoms within the past two weeks were queried with seven items, which are scored on a four-point Likert scale rated



from 0 (not at all) to 3 (nearly every day). The total GAD-7 score can range from 0 to 21. A score of 0 to 4 indicates the absence of generalized anxiety disorder, score of 5 to 9 represents mild, score of 10 to 14 represents moderate, and score of 15 and higher represents serve anxiety symptom levels (Löwe et al. 2008).

## Statistical analyses

We compared participants and non-participants in multiple logistic regression models according adjusted differences in age, sex, education, treatment setting, cancer type, and study center.

We calculated frequencies as well as means and standard deviations, as appropriate. Sociodemographic and medical groups were compared in terms of anxiety symptoms according to the GAD-7 mean score with one-way analysis of variance (ANOVA).

To dichotomize the scores and divide them into "anxious" and "non-anxious", the cut of  $\geq 10$  was chosen, following the recommendation of Andersen et al. (2014). To compare cancer patients with different types of cancer during and after treatment with the general population, we performed multivariate analyses using logistic regression. We adjusted odds ratios for confounders by including ages and sex in the regression model. Gender-specific tumors were the exceptions. Women with breast cancer and female genital tumors were compared only with women from general population, and patients with testicular cancer and prostate cancer were compared only with men. An adjustment was made here according to age. A p value < 0.05 was considered statistically significant. Analyses were performed using SPSS version 27 (SPSS Inc., Chicago, III, USA).

#### Results

# Characteristics of the sample of cancer patients

Out of 5,889 eligible cancer patients, 4,020 (69%) participated. Participants in this group were younger (p < 0.001), more educated (p < 0.001) and were more likely to be recruited from a cancer rehabilitation center (p < 0.001) than during treatment from acute care hospitals. We found no sex difference in study participation (p < 0.10). Participants' mean age was 58 years (SD=11). Mean time since diagnosis was 13.5 months (SD=25). Demographic and medical sample characteristics are presented in Table 1.

#### **Anxiety in cancer patients**

Out of all cancer patients, 13.8% showed elevated anxiety (GAD  $\geq$  10), while severe anxiety (GAD  $\geq$  15) were

relatively rare across tumor entities (Fig. 1). A significantly greater proportion of women compared with men reported anxiety (16.8 vs. 10.7%, p < 0.001). The highest prevalence of anxiety was observed in patients in the age category 36–45 years (20.1%), in single patients (13.2%), or in unemployed patients (21.3%). Even if not significant, a trend toward a greater proportion of patients in rehabilitation reported anxiety compared with patients during treatment (15.9 vs. 13.3% resp. 13.0%, p = 0.102). Nevertheless, the mean expression of anxiety is significantly greater in cancer rehabilitation than during treatment (5.5 vs. 5.1 resp. 5.0, p = 0.013). Furthermore, regarding the type of therapy, patients reported more anxiety after chemotherapy. The prevalence of anxiety was most evident in patients with testicular cancer, followed by bladder cancer, female genital tumors, patients with soft tissue tumors, lung cancer, and thyroid cancer. Patients with prostate carcinoma or pancreatic carcinoma reported relatively little anxiety. All results are listed in Table 1.

# Anxiety in cancer patients compared to general population

The two samples do not differ significantly in terms of age and sex. The proportion of anxiety in cancer patients is significantly higher at 13.8% compared to 5.7% in the general population (p < 0.001). Multivariate logistic regression analyses adjusted for age and sex showed significantly higher odds for anxiety in cancer patients (OR, 2.7; 95% CI 2.4–3.1; p < 0.001). We found similar results for the different treatment settings. Thus, the risk of anxiety was significantly higher in patients during cancer rehabilitation than in the general population (OR, 3.1; 95% CI 2.6 - 3.8; p < 0.001). It was increased 2.8-fold (95% CI 2.3–3.3; p < 0.001) during inpatient treatment and 2.4-fold (95% CI 2.0–2.9; p < 0.001) during outpatient treatment compared to the general population. An increased risk for anxiety was revealed in all other tumor entities except for patients with pancreatic carcinoma. Particularly, patients with bladder cancer were most likely to report anxiety with odds more than 5 times higher than those in general population (OR, 5.3; 95% CI 3.0-9.4). We observed a significantly increased risk for anxiety in testicular cancer patients (OR, 5.0; 95% CI 2.1-12.1). All results are listed in Table 2.

## Discussion

In this study, we aimed to measure and analyze the prevalence of anxiety in cancer patients taking into account sociodemographic and medical variables. We further compared levels of anxiety among cancer patients to the general population.



**Table 1** Sociodemographic and medical sample characteristics of anxiety (GAD-7) in cancer patients in Germany

|                                       | Total sample   |      | Anxiety (GAD-7≥10) |      | GAD-7          |      |         |
|---------------------------------------|----------------|------|--------------------|------|----------------|------|---------|
|                                       | $\overline{N}$ | %    | $\overline{N}$     | %    | $\overline{M}$ | SD   | p       |
| Sex                                   | ,              | ,    |                    |      |                |      |         |
| Woman                                 | 2068           | 51.4 | 329                | 16.8 | 5.70           | 4.16 | < 0.001 |
| Men                                   | 1952           | 48.6 | 194                | 10.7 | 4.57           | 3.87 |         |
| Age category <sup>a</sup>             |                |      |                    |      |                |      |         |
| 18–35                                 | 151            | 3.8  | 26                 | 18.4 | 5.6            | 4.02 | < 0.001 |
| 36–45                                 | 385            | 9.7  | 74                 | 20.1 | 6.0            | 4.34 |         |
| 46–55                                 | 897            | 22.5 | 157                | 18.2 | 5.9            | 4.23 |         |
| 56–65                                 | 1265           | 31.8 | 151                | 12.8 | 5.0            | 4.05 |         |
| 66–75                                 | 1286           | 32.3 | 110                | 9.2  | 4.4            | 3.71 |         |
| In a relationship <sup>a</sup>        |                |      |                    |      |                |      |         |
| Yes                                   | 2904           | 80.7 | 377                | 13.2 | 5.1            | 4.22 | 0.021   |
| No                                    | 696            | 19.3 | 113                | 16.7 | 5.5            | 4.02 |         |
| Education                             |                |      |                    |      |                |      |         |
| ≤10 years                             | 2456           | 61.1 | 343                | 14.3 | 5.2            | 4.16 | 0.459   |
| > 10 years                            | 1564           | 38.9 | 180                | 13.8 | 5.1            | 3.89 |         |
| Work situation <sup>a</sup>           |                |      |                    |      |                |      |         |
| Employed                              | 1569           | 42.5 | 223                | 14.1 | 5.3            | 3.99 | < 0.001 |
| Unemployed                            | 208            | 5.6  | 44                 | 21.3 | 6.1            | 3.99 |         |
| Retired                               | 1732           | 46.9 | 214                | 12.7 | 4.9            | 4.11 |         |
| Housewife/househusband                | 187            | 5.1  | 20                 | 11.0 | 4.7            | 3.75 |         |
| Cancer care setting                   |                |      |                    |      |                |      |         |
| Inpatients oncology ward              | 1735           | 43.2 | 213                | 13.3 | 5.1            | 4.04 | 0.013   |
| Outpatient oncology clinics           | 1324           | 32.9 | 162                | 13.0 | 5.0            | 3.96 |         |
| Cancer rehabilitation                 | 961            | 23.9 | 148                | 15.9 | 5.5            | 4.23 |         |
| Cancer type                           |                |      |                    |      |                |      |         |
| Testis                                | 36             | 0.9  | 7                  | 20.6 | 5.4            | 4.53 | < 0.001 |
| Bladder                               | 90             | 2.2  | 16                 | 19.8 | 5.4            | 4.16 |         |
| Female genital organs                 | 317            | 7.9  | 53                 | 17.5 | 5.9            | 4.17 |         |
| Soft tissue                           | 39             | 1.0  | 6                  | 17.1 | 6.2            | 4.51 |         |
| Lung                                  | 331            | 8.2  | 49                 | 16.8 | 5.2            | 4.41 |         |
| Thyroid                               | 25             | 0.6  | 4                  | 16.7 | 5.6            | 3.88 |         |
| Breast                                | 906            | 22.5 | 140                | 16.2 | 5.5            | 4.08 |         |
| Head and neck                         | 127            | 3.2  | 18                 | 15.4 | 4.9            | 4.20 |         |
| Stomach/esophagus                     | 146            | 3.6  | 21                 | 14.9 | 5.6            | 4.40 |         |
| Brain                                 | 62             | 1.5  | 8                  | 13.8 | 5.3            | 4.01 |         |
| Hematological                         | 305            | 7.6  | 40                 | 13.7 | 5.4            | 4.04 |         |
| Malignant melanoma                    | 67             | 1.7  | 9                  | 13.4 | 5.5            | 4.36 |         |
| Colon/rectum                          | 510            | 12.7 | 59                 | 12.4 | 4.9            | 3.88 |         |
| Kidney/urinary tract                  | 131            | 3.3  | 15                 | 12.3 | 4.7            | 4.35 |         |
| Hepatobiliary                         | 52             | 1.3  | 6                  | 11.8 | 5.0            | 4.16 |         |
| Prostate                              | 637            | 15.8 | 48                 | 8.1  | 4.1            | 3.54 |         |
| Pancreas                              | 82             | 2.0  | 6                  | 7.7  | 4.8            | 3.46 |         |
| Other                                 | 157            | 3.9  | 18                 | 12.2 | 5.2            | 4.13 |         |
| Cancer disease condition <sup>a</sup> | 137            | 3.7  | 10                 | 12.2 | 3.2            |      |         |
| In remission                          | 1539           | 39.4 | 200                | 13.6 | 5.1            | 3.98 | 0.683   |
| Not in remission                      | 2365           | 60.6 | 310                | 14.1 | 5.2            | 4.12 | 0.000   |



Table 1 (continued)

|                                      | Total sample |      | Anxiety<br>(GAD-7≥10) |      | GAD-7          |      |         |
|--------------------------------------|--------------|------|-----------------------|------|----------------|------|---------|
|                                      | N            | %    | $\overline{N}$        | %    | $\overline{M}$ | SD   | p       |
| Tumor stage /UICC TNM) <sup>a</sup>  |              |      |                       |      |                |      |         |
| I                                    | 539          | 13.6 | 71                    | 13.7 | 5.2            | 3.89 | 0.459   |
| II                                   | 700          | 17.7 | 93                    | 14.2 | 5.1            | 4.19 |         |
| III                                  | 546          | 13.8 | 63                    | 12.2 | 4.8            | 3.80 |         |
| IV                                   | 889          | 22.4 | 124                   | 15.2 | 5.2            | 4.19 |         |
| Metastases <sup>a</sup>              |              |      |                       |      |                |      |         |
| Yes                                  | 870          | 21.9 | 119                   | 14.9 | 5.3            | 4.19 | 0.491   |
| No                                   | 2400         | 60.4 | 298                   | 13.2 | 5.1            | 3.98 |         |
| Treatment intention <sup>a</sup>     |              |      |                       |      |                |      |         |
| Curative                             | 2396         | 59.6 | 292                   | 12.8 | 5.0            | 3.95 | 0.008   |
| Palliative                           | 926          | 23.0 | 140                   | 16.4 | 5.6            | 4.35 |         |
| Surgery <sup>a</sup>                 |              |      |                       |      |                |      |         |
| Yes                                  | 1039         | 26.8 | 388                   | 14.4 | 5.2            | 4.10 | 0.065   |
| No                                   | 2838         | 73.2 | 121                   | 12.6 | 4.9            | 3.98 |         |
| Radiation <sup>a</sup>               |              |      |                       |      |                |      |         |
| Yes                                  | 1762         | 45.1 | 239                   | 14.2 | 5.1            | 4.09 | 0.949   |
| No                                   | 2145         | 54.9 | 272                   | 13.6 | 5.1            | 4.05 |         |
| Chemotherapy <sup>a</sup>            |              |      |                       |      |                |      |         |
| Yes                                  | 2023         | 51.6 | 290                   | 15.3 | 5.3            | 4.17 | 0.001   |
| No                                   | 1896         | 48.4 | 221                   | 12.4 | 4.9            | 3.93 |         |
| Hormone therapy <sup>a</sup>         |              |      |                       |      |                |      |         |
| Yes                                  | 481          | 87.7 | 67                    | 14.5 | 5.2            | 3.95 | 0.691   |
| No                                   | 3420         | 12.3 | 444                   | 13.8 | 5.1            | 4.09 |         |
| ECOG performance status <sup>a</sup> |              |      |                       |      |                |      |         |
| 0                                    | 1900         | 49.2 | 200                   | 11.1 | 4.7            | 3.83 | < 0.001 |
| 1                                    | 1368         | 35.4 | 218                   | 16.8 | 5.5            | 4.24 |         |
| 2                                    | 465          | 12.0 | 63                    | 14.8 | 5.5            | 4.02 |         |
| 3                                    | 113          | 2.9  | 20                    | 21.3 | 6.4            | 4.59 |         |
| 4                                    | 13           | 0.3  | 3                     | 25.0 | 6.3            | 3.44 |         |

GAD-7, Generalized Anxiety Disorder–German version, N number of patients, M mean, SD standard deviation, \*Reduced sample size due to missing data, p value based on one-way ANOVA

# **Anxiety in cancer patients**

In our study, anxiety was found in almost 14% of cancer patients. This result is similar to the results of a cross-sectional study in which data on anxiety were collected with the Brief Symptom Inventory. The prevalence of anxiety in outpatients was about 12% (Brintzenhofe-Szoc et al. 2009). Another study, also surveyed self-reported anxiety, reported more frequently anxiety (22%) (Van Den Brekel et al. 2020). Because this study surveyed only oncology patients at the time of hospital admission, uncertainty about the subsequent course and upcoming treatments could be an indication of higher level of anxiety. However, this difference remains even when anxiety levels are considered with our inpatients (13%).

Another study with a large number of participants reported 19% of cancer patients showed clinical level of self-reported anxiety (Linden et al. 2012). Nevertheless, the measurement instruments used were different. The survey time points are also not comparable. For example, Linden et al. (2012) collected data before the first visit to a cancer center and thus before the start of treatment. In Brazil, outpatient oncology patients were found to have anxiety symptoms in 21.4%, also measured by the GAD-7 (Polidoro Lima and Osório 2014). Nevertheless, comparability with our study results is limited because the authors did not reported cut-off values used. Our results included samples from different treatment settings. However, the frequencies for anxiety did not differ between during treatment (inpatient and outpatient), and in rehabilitation. In contrast, however, a higher level of anxiety was observed after the



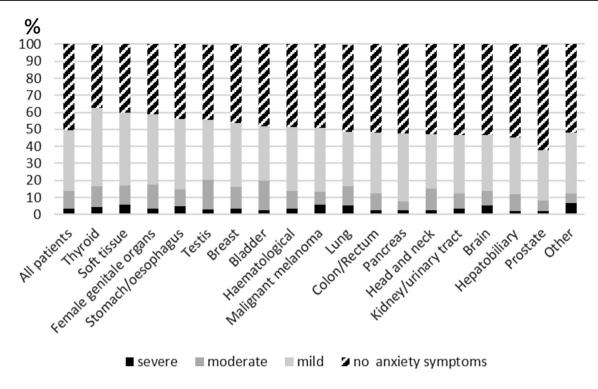


Fig. 1 Severity of anxiety (GAD-7 scores) by tumor entities in Germany. GAD-7 Generalized Anxiety Disorder-German version

end of therapy. Presumably, anxiety level is high before the start of treatment, because patients do not yet know what they will have to face. Anxiety after completing treatment could also include fear of disease recurrence. Yi and Syrjala (2017) stated, that patients after treatment report feeling alone or even abandoned following the intensive support provided during their treatment, and survivors often anxious and hypervigilant to physical sensations. A study from Germany observed that 9% of survivors reported moderate to severe anxiety 5 years after their cancer (Götze et al. 2020).

In our study, women reported anxiety more frequently and at higher levels than men. These findings generally fit into the literature that women reported more anxiety (Linden et al. 2012, Van Den Brekel et al. 2020) and distress related to cancer (Carlson et al. 2019; Herschbach et al. 2020). It seems that men focus more on physical problems while women tend to report emotional concerns (Loscalzo et al. 2018). We observed the highest proportion of anxiety in patients in the age group between 36 and 45 years. We suspect that this could be related to the current phase of life. Possibly the focus here is on anxiety about small children and the family. With increasing age, the proportion of patients who reported anxiety decreased. This finding is consistent with the results of the study by Meier et al. (2020) in which patients with hematologic cancers aged 70 and older reported low mean anxiety.

Both Linden et al. (2012) and Polidoro Lima and Osório (2014) demonstrated that anxiety was highest in women with

gynecologic tumors. In our sample, patients with testicular tumor, bladder cancer, and soft tissue tumors had the greatest anxiety. However, these entities were not included in the sample in the two studies cited. Following this, women with gynecologic tumors also had very high anxiety in our sample. In contrast, others found increased anxiety in patients with lymphoma as well as breast cancer (Brintzenhofe-Szoc et al. 2009). These patient groups also reported a high level of anxiety in our study.

# Anxiety in cancer patients compared to general population

Comparison of the prevalence of anxiety in cancer patients with the general population in Germany yielded a 2.7-fold increased risk for anxiety in cancer patients. With a 3.1-fold increased risk, patients in rehabilitation are even higher. We suspect that at this point the fear of late effects of the cancer and recurrence of the disease may increase. Perhaps patients who have completed treatment will become aware of their situation during rehabilitation. Even 5 years after the disease, a significantly higher level of anxiety, measured with GAD-7, was observed in cancer survivors in the age group up to 60 years in Germany compared to the general population (Götze et al. 2020).

Most of the tumor entities we studied are also above the general risk. Patients with testicular or lung cancer and, above all, bladder cancer were clearly at higher risk. These



**Table 2** Odds ratios for risk for anxiety in cancer patients (N=4.020) compared to general population sample (N=10.000) in Germany

|                            | Anxiety (GAD-7≥10) |       |      |         |  |  |
|----------------------------|--------------------|-------|------|---------|--|--|
|                            | OR                 | 95% ( | CI   | p       |  |  |
|                            |                    | LB    | UB   |         |  |  |
| All cancer patients        | 2.7 <sup>a</sup>   | 2.4   | 3.1  | < 0.001 |  |  |
| Setting                    |                    |       |      |         |  |  |
| Cancer rehabilitation      | $3.1^{a}$          | 2.6   | 3.8  | < 0.001 |  |  |
| Inpatient oncology wards   | $2.8^{a}$          | 2.3   | 3.3  | < 0.001 |  |  |
| Outpatient oncology clinic | $2.4^{a}$          | 2.0   | 2.9  | < 0.001 |  |  |
| Cancer type                |                    |       |      |         |  |  |
| Bladder                    | 5.3 <sup>a</sup>   | 3.0   | 9.4  | < 0.001 |  |  |
| Testis                     | $5.0^{b}$          | 2.1   | 12.1 | < 0.001 |  |  |
| Lung                       | $3.9^{a}$          | 2.8   | 5.4  | < 0.001 |  |  |
| Soft tissue                | $3.6^{a}$          | 1.5   | 8.9  | 0.004   |  |  |
| Head and neck              | $3.5^{a}$          | 2.1   | 5.9  | < 0.001 |  |  |
| Stomach/ esophagus         | $3.5^{a}$          | 2.2   | 5.6  | < 0.001 |  |  |
| Thyroid                    | $3.1^{a}$          | 1.0   | 9.1  | 0.043   |  |  |
| Malignant melanoma         | $2.8^{a}$          | 1.3   | 5.6  | 0.005   |  |  |
| Female genital organ       | 2.7 <sup>c</sup>   | 2.0   | 3.7  | < 0.001 |  |  |
| Brain                      | $2.6^{a}$          | 1.2   | 5.5  | 0.015   |  |  |
| Kidney/urinary tract       | $2.6^{a}$          | 1.5   | 4.6  | 0.001   |  |  |
| Hematological              | $2.6^{a}$          | 1.8   | 3.7  | < 0.001 |  |  |
| Colon/ rectum              | $2.6^{a}$          | 1.9   | 3.4  | < 0.001 |  |  |
| Breast                     | 2.5 <sup>c</sup>   | 2.0   | 3.1  | < 0.001 |  |  |
| Hepatobiliary              | 2.5 <sup>a</sup>   | 1.0   | 5.9  | 0.038   |  |  |
| Prostate                   | 2.4 <sup>b</sup>   | 1.7   | 3.3  | < 0.001 |  |  |
| Other                      | $2.4^{a}$          | 1.5   | 4.0  | 0.001   |  |  |
| Pancreas                   | 1.5 <sup>a</sup>   | 0.6   | 3.5  | 0.342   |  |  |

Confounders adjusted for age and sex, bmen, adjusted for age, women adjusted for age

OR, odds ratio; GAD-7, Generalized Anxiety Disorder–German version, 95% CI 95% confidence interval, LB lower boundary, UB upper Boundary

results are partly in the line with other studies. A systematic review indicates that patients with lung cancer experienced significantly more symptom distress than other cancer patients (Cooley 2000). A study from Scotland of 170 patients with lung cancer identified also an increased self-rated anxiety in over half of the patients (Buchanan et al. 2010). However, only patients in a palliative situation were included there. Regarding the association of bladder cancer and anxiety, another review showed very different results. The prevalence was 9.8% before treatment and ranged from 12.5 to 23% after treatment in Europe (Vartolomei et al. 2018).

Compared with the general population, patients with prostate cancer reported the least anxiety. One explanation could lie in the gender specificity of this condition, as we were able to show that men generally reported less anxiety than women did (Loscalzo and Clark 2018). Sánchez Sánchez et al. (2020) reported a prevalence of anxiety of 14.1% in 184 patients with prostate cancer as measured by the Hospital Anxiety and Depression Scale in Spain. In this regard, higher level of anxiety also appeared to be associated with multimodal treatment.

Surprisingly, we did not find a significant difference in patients with pancreatic cancer. A relationship with depressiveness does not seem to exist here, then this group of patients had a 7.8-fold increased risk of developing depression (Hartung et al. 2017). With a prevalence of anxiety of 7.7%, these patients are clearly below the prevalence of cancer patients in general in our analysis. A statistically significant difference to the general population could not be observed. Again, this could also be due to the small sample size. At this point, our results are contradictory to results of other studies that showed more anxiety in patients with pancreatic cancer (Kenner 2018). Certainly, social desirability of the responses, as well as different tumor stages and inconsistent measurement instruments, could be decisive here as well and complicate the comparison.

# **Strengths and limitations**

Strengths of the study are the multi-center design, especially the inclusion of patients with different tumor entities with different settings. The participation rate of 69% can be considered very high. Another advantage of our analyses is the comparison with prevalence data for anxiety from a large German general population. An important criterion for this is the use of the same measurement instrument and the same cut-off value. Nevertheless, there are some sources of bias to discuss that could affect the generalizability of the results. The most common reason given for non-participation was "too burdened". Perhaps these patients in particular were very anxious, also of psychological stigmatization. One more limitation mainly to be seen is the cross-sectional study. Future studies should prefer a measurement over time as a longitudinal section. In our sample, we did not check how many or which patients had a second tumor. With respect to the sample of the general population, we also cannot make any statements regarding possible comorbidities and the level of education. Statements regarding comparisons in terms of tumor entities must also be viewed with caution, as the studies included very different samples.

# **Conclusion**

In our study, a high prevalence of anxiety was observed in cancer patients and a significantly higher risk compared to the general population. Since physicians and nurses very



often overlook the presence of clinically relevant anxiety in oncology patients and assume an appropriate emotional response, it is important to identify anxiety in cancer patients at an early stage. These are related to disease and treatment. Also after completion the treatment, an increased level of anxiety is observed. Anxiety can negatively affect treatment adherence and have a significant impact on quality of life.

**Author contributions** AM-T and UK-G contributed to the study conception and design. Data analysis were performed by AM-T, UG, AH, JMH, and Peter Esser. The first draft of the manuscript was written by Ute Goerling and all authors commented on previous versions of the manuscript. All authors read an approved the final manuscript.

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**Data availability** The datasets generated during and/or analyzed during the current study are available from the corresponding author on reasonable request.

#### **Declarations**

**Conflict of interest** The authors have non-relevant financial or non-financial interests to disclose.

Ethical approval This study was performed in line with the principles of the Declaration of Helsinki. Approval was granted by the Ethics Committees in all involved federal states (Hamburg: Ref. Nr. 2768, Schleswig–Holstein: Ref. Nr. 61/09, Freiburg: Ref. Nr. 244/07, Heidelberg: Ref. Nr. S-228/2007; 50155039; Würzburg: Ref. Nr. 107/07; Leipzug: Ref. Nr. 200–2007).

Consent to participate Informed consent was obtained from all individual participants included in the study.

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