Radboud University Nijmegen

# PDF hosted at the Radboud Repository of the Radboud University Nijmegen

The following full text is a publisher's version.

For additional information about this publication click this link. http://hdl.handle.net/2066/207398

Please be advised that this information was generated on 2020-01-01 and may be subject to change.

# **RESEARCH ARTICLE**

# **Open Access**



# Predictors for use of psychosocial services in patients with metastatic colorectal cancer receiving first line systemic treatment

Claudia S. E. W. Schuurhuizen<sup>1,2</sup>, Annemarie M. J. Braamse<sup>3</sup>, Inge R. H. M. Konings<sup>1</sup>, Henk M. W. Verheul<sup>1†</sup> and Joost Dekker<sup>2\*†</sup>

# Abstract

**Background:** Patients with advanced disease experience high levels of psychological distress, yet there is low uptake of psychosocial services offered to patients who screened positive for distress. In this study we aimed to identify predictors for use of psychosocial services in patients with metastatic colorectal cancer (mCRC) receiving first line chemotherapy enrolled in a prospective cluster randomized trial (CRT).

**Methods:** Patients completed measures on psychological distress, physical distress, and quality of life at baseline. Demographics, clinical characteristics at baseline and clinical events during treatment (e.g. severe adverse events, clinical benefit) were extracted from patient records. Patients reported psychosocial service utilization in- and outside the hospital after 10, 24 and 48 weeks of treatment. Multivariable logistic regression models were used to identify predictors for the use of psychosocial services.

**Results:** Out of 349 patients, seventy patients (20.0%) used psychosocial support services during the follow-up period. Use of psychosocial services was associated with younger age, a higher educational level, presence of more pain (at baseline), and the expressed need to talk to a professional (at baseline). In addition, patients without progressive disease within the first ten weeks of treatment were more likely to use psychosocial services .

**Conclusions:** One in five patients with mCRC receiving first line palliative treatment used psychosocial services during this prospective longitudinal CRT. Sociodemographic factors (age, education), clinical factors (pain and no progressive disease) and the expressed need to talk to a professional predicted use of psychosocial services. Identification of these predictors may contribute to the understanding of factors that determine the need for psychosocial services.

Trial registration: Netherlands Trial Register NTR4034.

Keywords: Metastatic colorectal cancer, Palliative treatment, Psychosocial services, Predictors

\* Correspondence: j.dekker@vumc.nl

<sup>+</sup>Henk M. W. Verheul and Joost Dekker contributed equally to this work. <sup>2</sup>Department of Psychiatry and Amsterdam Public Health research institute, VU University Medical Center, Amsterdam, the Netherlands Full list of author information is available at the end of the article



© The Author(s). 2019 **Open Access** This article is distributed under the terms of the Creative Commons Attribution 4.0 International License (http://creativecommons.org/licenses/by/4.0/), which permits unrestricted use, distribution, and reproduction in any medium, provided you give appropriate credit to the original author(s) and the source, provide a link to the Creative Commons license, and indicate if changes were made. The Creative Commons Public Domain Dedication waiver (http://creativecommons.org/publicdomain/zero/1.0/) applies to the data made available in this article, unless otherwise stated.

### Background

Over the past decades, research has demonstrated the considerable emotional impact that colorectal cancer (CRC) and its treatment can have on patients [1]. Self-reported psychological distress occurs in approximately one third of CRC patients [2-5]. Patients with advanced disease seem to be even more susceptible to develop symptoms of psychological distress [6]. A higher level of psychological distress is associated with lower quality of life (QOL), non-adherence to therapy, and poorer prognosis [7–9]. A variety of psychological interventions have been shown to be effective in reducing symptoms of distress and improving QOL [10]. As a result, current guidelines for cancer care have recommended to routinely screen for psychological distress and to subsequently offer treatment to those scoring above the cut off for distress [11].

Yet, the majority of patients scoring above the cutoff for distress decline professional intervention [12-15]. It seems that the presence of psychological distress does not necessarily equate to the need of patients for psychosocial support [2, 12, 16-18]. Distress as such does not seem to be a sufficient reason to use psychosocial services, as many patients scoring above the cut off for distress decline these services. Therefore, instead of assuming that all distressed patients need psychosocial services, it seems important to take one step back and to compare patients who use services against those who do not. Knowledge on factors related to the actual use of psychosocial services may contribute to our understanding of the factors that determine the need for psychosocial services. We aimed to identify predictors for use of psychosocial services in patients with mCRC receiving first line treatment. Potential predictors included demographic, clinical, and psychosocial factors at baseline and clinical factors during treatment.

# Methods

This paper reports on a secondary analysis on data obtained in the TES trial (Targeted selection, Enhanced care, Stepped care trial). In this trial the effectiveness of a combined screening and treatment program to improve psychological distress compared with care as usual was evaluated (Netherlands Trial Register (NTR4034) [19]. The study was approved by the Medical Ethics Committee of VU University Medical Center (2013/10). Analyses are based on 349 outpatients with a confirmed diagnosis of mCRC starting first line systemic treatment who met the eligibility criteria, recruited throughout 16 hospitals in the Netherlands, between July 2013 and October 2016. All patients provided written informed consent. A description of the study protocol and results of the original trial have been published previously [14, 19]. In brief, there were no significant differences in course of distress between the treatment arms. Further, no differences in recognition and management of distress or use of psychosocial support between groups were found, although satisfaction with care and cognitive functioning were significantly better in the intervention group.

#### Outcome and measurements

Demographic and clinical characteristics were obtained from patients and medical charts at baseline, i.e. start of first line systemic treatment. In addition, during the 48 weeks of the TES study, information on clinical events was collected. At baseline, patient-reported psychological characteristics were measured with the following tools. The HADS (Hospital Anxiety and Depression Scale) and DT/PL (Distress Thermometer and corresponding Problem List) were used to assess distress and the need for referral with a recall period of two weeks, respectively. The HADS consist of 14 questions with scores ranging from 0 to 42, the recommended cut off score for distress in cancer patients is  $\geq 13$  [20]. The DT/PL comprises of the Distress Thermometer ranging from 0 to 10, followed by a list of problems evaluating a wide range of problems and a single question evaluating the need to talk to a professional [2]. A score of 5 or higher on the DT has been shown to indicate presence of distress [2]. QOL was measured by the EORTC-QLQ-C30 version 3.0 [21], consisting of five functional scales (physical, role, cognitive, emotional, and social), symptom scales and individual symptom items, and perceived financial impact of the disease.

During treatment, the following clinical events were assessed by clinicians: treatment modification or discontinuation, clinical benefit during the first 10 weeks of treatment (progression vs stable disease or partial response), occurrence of severe (i.e. grade III/IV) adverse events (AEs, these were graded and assigned by clinicians using the National Cancer Institute's Common Terminology Criteria for Adverse Events (NCI-CTCAE) as mild (Grade 1), moderate (Grade 2), severe (Grade 3), or life-threatening (Grade 4) [22]) and total number of unscheduled admissions. Psychosocial service utilization in- and outside the hospital was assessed using an adapted version of the TiC-P questionnaire [23] after 10, 24 and 48 weeks of treatment with a recall period of three months. The TiC-P is a feasible and reliable instrument for collecting data on medical consumption and productivity losses [23, 24]. The first part of the TiC-P includes 14 structured yes/no questions on relevant medical resource items each followed by a question on the volume of medical consumption. The questions include contacts within the mental healthcare sector (regional mental healthcare organization, psychiatrist/ psychologist or psychotherapist in private practices or outpatient hospital, institutional day-care treatment,

Consultation Agency for Alcohol and Drug addiction (CAD), self-help group) and contacts with general healthcare providers (general practitioner, allied health professional, social worker, providers of alternative medicine, outpatients visits to medical specialists, hospital admission and contacts with a company doctor) and the use of medication. For the current analyses we evaluated contacts with social workers, contacts with services in regional mental healthcare organizations, contacts with psychiatrists/psychologists in private practice, and contacts with psychiatrists/psychologist in hospital settings.

#### Statistical analyses

Descriptive statistics were calculated to summarize all variables and to outline the pattern of use of psychosocial services in patients from the TES trial. Differences between patients who had used psychosocial services versus those who had not used psychosocial services during follow-up were computed using the chi-square test for categorical data and t-test for metric data. Two separate multivariable analyses were undertaken to identify predictors for the use of psychosocial support. In the first analysis, baseline variables potentially related to the use of psychosocial support were evaluated. Explanatory variables included gender (male vs female), age at randomization, married/living together vs living alone, education level (low: primary education, middle: lower general secondary education, intermediate vocational education or high school, high: higher vocational education or university), employment (working yes vs no), functional status (defined by the Eastern Cooperative Oncology Group Performance Status (ECOG PS), which is a score ranging from zero ("fully active") through three ("capable of only limited self-care") to five ("dead") [25]), right or left sided tumor, number of prior cancer-related treatments, type of chemotherapy (capecitabine vs CAPOX vs FOLFOX or other), total number of (organs with) metastases, time since initial diagnosis in months (< 1.5 months vs 1.5-10 months vs > 10 months), time since diagnosis of metastatic disease in months (< 1.0 months vs 1.0–2.5 months vs > 2.5 months), comorbidity (number of comorbid conditions as reported by patients at baseline), allocation to treatment arm of the parent study (intervention vs control) and participation in other clinical studies (yes vs no), distress on HADS (score of <13 or  $\ge$  13), distress score on DT (< 5 or  $\ge$  5), number of physical problems reported on PL, number of emotional problems reported on PL, need to talk to a professional (yes vs no), scores on functional QOL scales, global QOL, fatigue and pain score on EORTC-QLQ-C30. Variables were included in the multivariable logistic regression equation if they had an univariable association  $(p \le .25)$ with the criterion [26]. A backward stepwise regression model was computed to select the final model that included predictive factors which were statistically significant at the 0.10 level. Variables not significantly contributing to the multivariable logistic regression equation were removed. In the second analysis it was evaluated whether clinical factors during treatment predict use of psychosocial support. Explanatory variables included clinical benefit during the first 10 weeks of chemotherapy treatment, number of severe AEs, any severe AE during treatment, any admission during treatment, and number of admissions. Using the final model on baseline predictors as a base, variables were included in this model if they had an univariable association ( $p \le .25$ ) with the criterion. A backward stepwise regression model was computed to select the final model that included predictive factors which were statistically significant at the 0.10 level. Variables not significantly contributing to the multivariable logistic regression equation were removed. The explained variance of the model was assessed by use of Nagelkerke's  $R^2$ . Since multicollinearity can influence the estimated beta parameter, the variable inflation factor (VIF) for each variable included in the regression equations was calculated (i.e. a VIF greater than 2.5 indicates multicollinearity) [27, 28]. Data were analyzed using IBM SPSS statistics version 22.0 (IBM Corp., Armonk, NY).

#### Results

The sample consisted of all 349 patients enrolled in the TES study [14]. Table 1 contains sample characteristics. The 349 patients ranged in age from 23 to 83 years at diagnosis (mean age  $66.1 \pm 10.2$  years); 224 (64.2%) were male, 256 (73.4%) were married/living with a partner, 93 (26.7%) scored above the cutoff for distress (measured with the HADS), and 34 (9.7%) expressed a need to talk to a professional. At the time of baseline assessment all patients were receiving chemotherapy, 72 patients (20.6%) monotherapy, 274 (78.5%) an oxaliplatin-based combination regimen; for 3 patients type of chemotherapy was unknown. All treatments were administered with palliative intent. Seventy (20.0%) out of 349 patients made use of psychosocial services during the 48-week follow-up period. Of these, 36 (51.4%) had contact with social workers, 6 (8.6%) had contact with services in regional mental healthcare organizations, 13 (18.6%) had contact with psychiatrists/psychologists in private practices, 4 (5.7%) had contact with psychiatrists/psychologists in the hospital setting and the remaining 27 patients (38.6%) had contact with multiple psychosocial services. Twenty-six of 93 (28.4%) patients who scored above the cutoff for distress on the HADS at baseline used psychosocial services during the 48-week follow-up period. Twelve of 34 (35.3%) patients who had expressed a need to talk to a professional, used psychosocial services during the 48-week follow-up period.

Table 1 contains the univariable analyses for baseline sociodemographic, clinical and psychological factors and

# Table 1 Results of univariable analyses on demographic, clinical and psychosocial baseline variables

	No use of psychosocial support services ( $n = 279$ )(n,%)	Use of psychosocial support services $(n = 70)(n,\%)$	p-value
Age, years			*0.003
Mean age (SD)	66.92 (9.7)	62.74 (11.41)	
Gender			0.415
Male	182 (65.2)	42 (60.0)	
Female	97 (34.8)	28 (40.0)	
Treatment arm in TES trial			0.610
Intervention	149 (53.4)	35 (50)	
Control	130 (46.6)	35 (50)	
ECOG PS <sup>a</sup>			0.714
0	67 (24.0)	15 (21.4)	
1	87 (31.2)	18 (25.7)	
2	8 (2.9)	3 (4.3)	
Missing	117 (41.9)	34 (48.6)	
Primary tumor location			0.695
Right-sided	84 (30.1)	23 (32.9)	
Left-sided	192 (68.8)	47 (67.1)	
Missing	3 (1.1)	-	
Chemotherapy regimen			0.875
Capecitabine	59 (21.1)	13 (18.6)	
CAPOX	190 (68.1)	50 (71.4)	
FOLFOX/Other	27 (9.7)	7 (10.0)	
Missing	3 (1.1)	-	
Number of comorbidities			0.980
Mean (SD)	2.33 (1.78)	2.33 (2.00)	
Marital status			0.369
Married/domestic partnership	207 (74.2)	49 (70.0)	
Unmarried/divorced/widowed	68 (24.4)	21 (30.0)	
Missing	4 (1.4)	-	
Number of persons in household			0.615
1	49 (17.6)	13 (18.6)	
2	177 (63.4)	42 (60.0)	
3	23 (8.2)	4 (5.7)	
> 3	26 (9.3)	11 (15.7)	
Missing	4 (1.4)	-	
Education <sup>b</sup>			*0.041
Low	17 (6.1)	2 (2.9)	
Middle	184 (65.9)	39 (55.7)	
High	72 (25.8)	29 (41.4)	
Missing	6 (2.2)	0 (0)	
Currently working			0.949
Yes	65 (23.3)	16 (22.9)	
No/retired	210 (75.3)	54 (77.1)	
Missing	4 (1.4)	-	
Time from diagnosis primary tumor until start study			0.848
< 1.5 months	94 (33.7)	22 (31.4)	
1.5-10 months	89 (31.9)	25 (35.7)	
	\ /	( /	

## Table 1 Results of univariable analyses on demographic, clinical and psychosocial baseline variables (Continued)

	No use of psychosocial support services ( $n = 279$ )( $n$ ,%)	Use of psychosocial support services $(n = 70)(n,\%)$	p-value
> 10 months	93 (33.3)	23 (32.9)	
Missing	3 (1.1)	_	
Time from diagnosis metastatic disease until start study			0.666
< 1.0 months	95 (34.1)	27 (38.6)	
1.0-2.5 months	87 (31.2)	23 (32.9)	
> 2.5 months	94 (33.7)	20 (28.6)	
Missing	3 (1.1)	_	
Prior cancer related treatment			
No	85 (30.5)	23 (32.9)	0.699
Yes	194 (69.5	47 (67.1)	
Prior treatment metastases			
No	214 (76.7)	56 (80.0)	0.657
Yes	62 (22.2)	14 (20.0)	
Missing	3 (1.1)	_	
Number of organs with metastases			0.599
Median	2	2	
Total number of metastases			
Median	11.5	13.0	0.877
Participation in other trials			0.718
No	170 (60.9)	41 (58.6)	
Yes	109 (39.1)	29 (41.4)	
HADS			
Total score	9.10 (6.30)	11.17 (7.47)	*0.020
Distress on HADS No	208 (74.6)	44 (62.9)	*0.033
Yes	67 (24.0)	26 (37.1)	
Missing	4 (1.4)	_	
DT/PL			
DT score	4.33 (2.68)	4.75 (2.40)	*0.242
Total of physical problems	5.78 (6.29)	7.37 (4.61)	*0.020
Total of emotional problems	1.95 (2.11)	2.74 (2.57)	*0.009
Need to talk to professional No	257 (92.1)	58 (82.9)	*0.023
Yes	22 (7.9)	12 (17.1)	
Quality of life (QLQ-C30)			
Physical functioning	74.84 (20.30)	72.76 (21.88)	0.451
Role functioning	65.69 (30.70)	58.81 (30.13)	*0.010
Emotional functioning	78.25 (18.12)	72.86 (21.41)	*0.095
Cognitive functioning	89.44 (14.97)	86.90 (16.76)	*0.219
Social functioning	78.08 (23.89)	69.05 (31.38)	*0.010
Global QOL	63.77 (21.73)	60.00 (21.78)	*0.197
Pain score	16.00 (23.75)	40.15 (26.57)	*0.174
Fatigue score	25.00 (16.67)	45.08 (44.44)	*0.009

Abbreviations: SD standard deviation, ECOG PS Eastern Cooperative Oncology Group Performance Status, HADS Hospital Anxiety and Depression Scale, DT/PL Distress Thermometer and Problem list, QLQ-C30 Quality of Life Questionnaire-C30. \*Characteristics were included in the multivariate model if they had an univariable association ( $p \le .25$ ). \*ECOG PS 0: Fully active, able to carry on all pre-disease performance without restriction. ECOG PS 1: Restricted in physically strenuous activity but ambulatory and able to carry out work of a light or sedentary nature, e.g., light house work, office work. ECOG PS 2: Ambulatory and capable of all selfcare but unable to carry out any work activities; up and about more than 50% of waking hours. <sup>b</sup>Low: primary education, middle: lower general secondary education, intermediate vocational education or high school, high: higher vocational education or university

their associations with the use of psychosocial support. Patients in the group that used psychosocial services were significantly younger at baseline and were more often higher educated. In addition, they reported higher levels of distress and were more frequently classified as being distressed (i.e. a score of  $\geq 13$  on HADS), reported more emotional and physical problems, and expressed the need to talk to a professional more frequently. Finally, patients who used psychosocial services demonstrated lower levels of role and social functioning at baseline and were significantly more fatigued. There were no differences for gender, marital status, employment, tumor burden or ECOG PS at baseline. Table 2 contains the univariable analyses for clinical events during follow-up and their associations with the use of psychosocial support. Patients in the group that used psychosocial services were more often responding to therapy within the first ten weeks of treatment (i.e. stable disease or partial response), whereas patients who did not use psychosocial services more often had had progression of disease within the first ten weeks of treatment.

Table 3 contains the regression parameters for the baseline predictors retained in the backward logistic regression analysis. Use of psychosocial services was associated with younger age, presence of more pain (at baseline), an expressed need to talk to a professional and a higher educational level. The final model with baseline characteristics had an overall classification rate of 81.2% with Nagelkerke  $R^2$  statistic of 0.118. In Table 4 the final model with baseline variables was used as a base and contains the regression parameters for the variables retained in the backward logistic regression analysis after adding clinical factors during treatment. Combined with the four baseline variables of the first model, patients who had progressive disease within the first ten weeks of treatment were less likely to use psychosocial services. The final model combining selected baseline variables and clinical variables over time had an overall classification rate of 81.5% with Nagelkerke  $R^2$  statistic of 0.140.

#### Discussion

This study prospectively investigated the predictors for use of psychosocial services in patients with mCRC receiving first line palliative treatment. Psychosocial support was broadly defined as contacts with social workers, contacts with services in regional mental healthcare organizations, contacts with psychiatrists/psychologists in private practice, and contacts with psychiatrists/psychologist in hospital settings. Our approach is unique as we assessed and distinguished possible predictors at baseline and clinical predictors during treatment. In line with previously reported results [16], approximately one in five patients used psychosocial services at any point during their 48-weeks clinical follow-up. In this sample of patients with mCRC, those patients having a younger age, having completed higher education levels, indicating higher levels of pain and expressing the need to talk to a professional were found to be more likely to use psychosocial services. Furthermore, absence of disease progression within the first ten weeks of treatment was found to be a clinical predictor for the use of these services.

The relationship between younger age and the use of psychosocial services is consistent with previous research [29-31]. A plausible explanation could be that advanced disease may be more traumatic and require greater life adjustments in younger patients who are in a different phase of life with work, children and relationships, compared to older patients [28, 31]. In addition, younger patients may be more aware of psychosocial services [28]. Users of psychosocial support in our study were more likely to have completed higher levels of education, a trend that is consistent with the literature [30, 32]. A knowledge gap may exist between patients with different education levels, those with less education may have greater information needs regarding the benefits and availability of psychosocial services [32]. In line with findings from O'Hea et al. [29], patients reporting more pain were seen to use psychosocial services more often. In agreement with our findings that showed that the

	No use of psychosocial support services (n = 279)(n,%)	Use of psychosocial support services (n = 70)(n,%)	p-value
Progression of disease within first 10 weeks of treatment			*0.089
No	242 (86.7)	66 (94.3)	
Yes	37 (13.3)	4 (5.7)	
Progression of disease during follow up			0.289
No	93 (33.3)	28 (40.0)	
Yes	186 (66.7)	42 (60.0)	
Total number of grade 3–4 AEs during 48 weeks follow up	1.64 (2.12)	2.06 (2.65)	*0.166
Total number of admissions during 48 weeks follow up	1.79 (1.34)	2.13 (1.98)	*0.216

Abbreviations: AEs adverse events. \*Characteristics were included in the multivariate model if they had an univariable association ( $p \le .25$ )

Predictive factor <sup>a</sup>	Wald	<i>p</i> -	Odds	95% C.I.for EXP(B)		VIF
		value	Ratio	Lower	Upper	
Need to talk to professional	4.451	.035	2.360	1.063	5.240	1.010
Pain score at baseline	8.053	.005	1.015	1.005	1.025	1.009
Age	8.066	.005	.963	.938	.988	1.022
Education	5.823	.054				1.031
Middle compared to low	.084	.772	1.255	.270	5.837	
High compared to low	1.340	.247	2.518	.527	12.024	

 Table 3 Final multivariable model of baseline predictors

Abbreviations: OR odds ratio, VIF variable inflation factor. <sup>a</sup>Predictive factors were statistically significant at the least at the 0.10 level

expressed need to talk to a professional was a predictor for use of psychosocial services, asking patients about their desire for help to highlight who are willing to accept psychosocial interventions has been proposed before as an efficient way of connecting patients to psychosocial services [16, 33].

The lack of association between gender and marital status and of use of psychosocial services contrasts with previous studies [28, 29, 31, 34], in which female and single patients have been shown to be using psychosocial services more often. However, those results were obtained mainly in heterogeneous study samples, predominated by younger, female patients suffering from breast cancer. Our study involved a homogeneous sample of on average older, mostly male mCRC patients, whose emotional concerns and needs, or access to psychosocial care services may have been different.

Absence of disease progression was associated with higher use of psychosocial services, while disease progression itself has previously been shown to be an important predictor for psychological distress [14, 35]. One hypothesis could be that patients' focus when facing disease progression may shift towards end-of-life issues, in which energy is more likely to be spent on preparation for death and achieving a sense of completion than in using psychosocial support services [36, 37].

Other clinical factors, such as burden of disease, time since diagnosis (of metastasis), type of systemic treatment, or amount of suffered high grade AEs and subsequent admissions did not appear to be predictors. Low predictive associations between clinical variables and need for psychological support services were demonstrated before [38]. These combined findings suggest that clinical (both pre- and on-treatment) variables are largely not predictive for use of psychosocial support, and emphasize that patient sociodemographic characteristics and self-report assessments of patients' physical and psychosocial well-being are essential in predicting which patients will use psychosocial services.

In line with previous work [34, 38], our results demonstrated that out of 93 patients who scored above the cutoff for distress at baseline, fewer than one third used psychosocial services. It has been demonstrated that the majority of patients show strong resilience following life-threatening events, characterized by transient symptoms of distress and return to normal levels without need for psychosocial interventions [39, 40]. This kind of emotional distress may even facilitate adaptation to cancer [12]. A minority of distressed patients does not seem to be able to cope with emotions related to cancer; this group of patients may be in need for referral to specialized mental healthcare [12].

Even though expressing the need to talk to professionals was found to be predictive for use of psychosocial services, out of 34 patients that expressed the subjective need to talk to a professional, services were only used by 12 (35.3%) of these patients. This latter finding is concerning. In our study we did not further

Table 4 Final multivariable model of baseline predictors and clinical predictors

Predictive factor <sup>a</sup>	Wald	<i>p-</i> value	Odds Ratio	95% C.I.for EXP(B)		VIF
				Lower	Upper	
Need to talk to professional	4.712	.030	2.429	1.090	5.412	1.010
Pain score at baseline	9.895	.002	1.017	1.006	1.028	1.030
Age	6.938	.008	.965	.940	.991	1.030
Education	6.190	.045				1.033
Middle compared to low	.128	.720	1.327	.283	6.229	
High compared to low	1.558	.212	2.729	.564	13.204	
Progression within first 10 weeks of treatment	4.136	.042	.314	.103	.959	1.029

Abbreviations: OR odds ratio, VIF variable inflation factor. <sup>a</sup>Predictive factors were statistically significant at the least at the 0.10 level

investigate why patients with an expressed need did or did not use support services. Known barriers to use of psychosocial services are lack of knowledge, financial constraints, lack of access, as well as lack of confidence in and negative perceptions towards psychosocial services [18, 41]. In order for those patients with a need for support to actually access these services, future studies should focus on the promotion of psychosocial support services [42]. However, it is worth mentioning that the item used to assess the need to talk to a professional could have been interpreted differently by patients, as there was no specific reference to mental health and this need may have referred to various disciplines. In a recently updated version of the Distress Thermometer, patients can now indicate with whom they would like to talk followed by a list of both physical and psychosocial healthcare providers [43].

#### **Study limitations**

Limitations of the present study include the self-reporting of use of psychosocial services, that may have under- or overestimated the actual use of these care services. Self-reported data are argued to be less reliable and threatened by bias arising from social desirability and recall period [44]. Second, this study did not make a clear distinction between type of psychosocial needs and problems. For some needs, especially needs resulting from practical or logistical issues, psychosocial services may not be appropriate. In addition, the analyses were conducted on a homogenous sample of particularly middle-aged, male, patients with advanced CRC participating in a CRT. Further research is needed to determine whether these findings apply to community-based settings as well and whether these findings are generalizable to younger or older patients. Furthermore, the need for psychosocial support was assessed with one single item question only. Another limitation was the sample size. Even though a total of 349 patients participated in the original trial, only 70 made use of psychosocial support. As a consequence some of the subgroups only contained a small number of patients, which could limit the generalizability of the results, warranting external validation.

#### **Clinical implications**

Our results can be used to inform clinicians in intervention practices as these provide insight into characteristics of patients receptive to psychosocial care. Identification of predictors for use of psychosocial services in patients with mCRC may allow resources to be directed more efficiently to patients who need and want help. In addition, our study findings underscore that relying on screening for distress to identify patients that use psychosocial services is less successful than has been assumed [33]: the need to talk to a professional, instead of the level of distress, predicted use of psychosocial services. Instead of relying on screening for distress, psychosocial services could be targeted to patients expressing the need to talk to a professional. This could be supplemented with more targeted assessments of patients at risk for psychosocial problems [45]. In addition, it should be emphasized that there may be a group of individuals at risk of suffering from negative psychosocial outcomes that is unlikely to use psychosocial services (e.g. less educated, older man), future research should focus on potential barriers to this use and on novel targeting tools to include these patients that my benefit from support.

# Conclusions

To conclude, approximately 20% of patients with mCRC starting with first line palliative treatment used psychosocial services during 48 weeks of follow up. The following predictors for use of psychosocial support were identified: being younger, reporting higher levels of pain, being higher educated, expressing a need to talk to a professional and the absence of progressive disease within the first weeks of treatment. These findings contribute to the understanding of factors that determine the need for psychosocial services.

#### Abbreviations

AEs: Adverse events; CAD: Consultation Agency for Alcohol and Drug addiction; CRC: Colorectal cancer; CRT: Cluster randomized controlled trial; DT: Distress Thermometer; DT/PL: Distress Thermometer / Problem List; ECOG PS: Eastern Cooperative Oncology Group Performance Status; HADS: Hospital Anxiety and Depression Scale; mCRC: metastatic colorectal cancer; NCI-CTCAE: National Cancer Institute's Common Terminology Criteria for Adverse Events; QOL: Quality of life; TES: Targeted selection, enhanced and stepped care; VIF: Variable inflation factor

#### Acknowledgements

An abstract of this paper has been presented at the 20th World Congress of Psycho-Oncology, 29 October - 2 November 2018 [46], Hong Kong: https://onlinelibrary.wiley.com/doi/full/10.1002/pon.4878

#### Funding

This work was supported by a grant from Dutch Cancer Society/Alpe d'Huzes (VU 2011–5279). The funding body had no role in the design of the study, collection, analysis, and interpretation of data or in writing the manuscript.

#### Availability of data and materials

The datasets used and/or analysed during the current study are available from the corresponding author on reasonable request.

#### Authors' contributions

CS: Conceptualization, data curation, methodology, formal analysis, writing – original draft and writing - review and editing. AB: Supervision, conceptualization, methodology, and writing - review and editing. IK: Supervision, conceptualization, and writing - review and editing. HV: Supervision, conceptualization, and writing - review and editing. JD: Supervision, conceptualization, methodology, and writing - review and editing. AD: Supervision, conceptualization, methodology, and writing - review and editing. AD: Supervision, conceptualization, methodology, and writing - review and editing. All authors have read and approved the manuscript.

#### Ethics approval and consent to participate

The study was approved by the Medical Ethics Committee of VU University Medical Center (2013/10). All patients provided written informed consent.

#### Consent for publication

Not applicable.

#### **Competing interests**

HV receives grants from Roche, Vitromics Healthcare System, Novartis, Immunovo B.V., Amgen, non-financial support from Pfizer, and honoraria from Boehringer Ingelheim. These resources were all received outside the submitted work. The other authors declare no competing interests.

#### **Publisher's Note**

Springer Nature remains neutral with regard to jurisdictional claims in published maps and institutional affiliations.

#### Author details

<sup>1</sup>Department of Medical Oncology, VU University medical center, Cancer Center Amsterdam, Amsterdam, the Netherlands. <sup>2</sup>Department of Psychiatry and Amsterdam Public Health research institute, VU University Medical Center, Amsterdam, the Netherlands. <sup>3</sup>Department of Medical Psychology and Amsterdam Public Health Institute, Academic Medical Center, Cancer Center Amsterdam, Amsterdam, the Netherlands.

#### Received: 2 October 2018 Accepted: 24 January 2019 Published online: 01 February 2019

#### References

- Zabora J, BrintzenhofeSzoc K, Curbow B, Hooker C, Piantadosi S. The prevalence of psychological distress by cancer site. Psychooncology. 2001; 10(1):19–28.
- Tuinman MA, Gazendam-Donofrio SM, Hoekstra-Weebers JE. Screening and referral for psychosocial distress in oncologic practice: use of the distress thermometer. Cancer. 2008;113(4):870–8.
- Carlson LE, Angen M, Cullum J, Goodey E, Koopmans J, Lamont L, MacRae JH, Martin M, Pelletier G, Robinson J, et al. High levels of untreated distress and fatigue in cancer patients. Br J Cancer. 2004;90(12):2297–304.
- Dunn J, Ng SK, Holland J, Aitken J, Youl P, Baade PD, Chambers SK. Trajectories of psychological distress after colorectal cancer. Psychooncology. 2013;22(8):1759–65.
- Linden W, Vodermaier A, Mackenzie R, Greig D. Anxiety and depression after cancer diagnosis: prevalence rates by cancer type, gender, and age. J Affect Disord. 2012;141(2–3):343–51.
- Diaz-Frutos D, Baca-Garcia E, Garcia-Foncillas J, Lopez-Castroman J. Predictors of psychological distress in advanced cancer patients under palliative treatments. Eur J Cancer Care (Engl). 2016;25(4):608–15.
- Brown KW, Levy AR, Rosberger Z, Edgar L. Psychological distress and cancer survival: a follow-up 10 years after diagnosis. Psychosom Med. 2003;65(4): 636–43.
- Quinten C, Coens C, Mauer M, Comte S, Sprangers MA, Cleeland C, Osoba D, Bjordal K, Bottomley A, Groups EC. Baseline quality of life as a prognostic indicator of survival: a meta-analysis of individual patient data from EORTC clinical trials. Lancet Oncol. 2009;10(9):865–71.
- DiMatteo MR, Lepper HS, Croghan TW. Depression is a risk factor for noncompliance with medical treatment: meta-analysis of the effects of anxiety and depression on patient adherence. Arch Intern Med. 2000; 160(14):2101–7.
- Faller H, Schuler M, Richard M, Heckl U, Weis J, Kuffner R. Effects of psychooncologic interventions on emotional distress and quality of life in adult patients with cancer: systematic review and meta-analysis. J Clin Oncol. 2013;31(6):782–93.
- Holland J, Watson M, Dunn J. The IPOS new international standard of quality Cancer care: integrating the psychosocial domain into routine care. Psychooncology. 2011;20(7):677–80.
- Dekker J, Braamse A, Schuurhuizen C, Beekman ATF, van Linde M, Sprangers MAG, Verheul HM. Distress in patients with cancer - on the need to distinguish between adaptive and maladaptive emotional responses. Acta Oncol. 2017;56(7):1026–9.
- Braamse AM, van Meijel B, Visser OJ, Boenink AD, Cuijpers P, Eeltink CE, Hoogendoorn AW, van Marwijk Kooy M, van Oppen P, Huijgens PC, et al. A randomized clinical trial on the effectiveness of an intervention to treat psychological distress and improve quality of life after autologous stem cell transplantation. Ann Hematol. 2016;95(1):105–14.

- 14. Schuurhuizen CS, Braamse AM, Beekman AT, Cuijpers P, van der Linden MH, Hoogendoorn AW: Screening and stepped care targeting psychological distress in patients with metastatic colorectal cancer: the TES cluster randomized trial: Accepted for publication pending minor revisions JNCCN—Journal of the National Comprehensive Cancer Network.
- Funk R, Cisneros C, Williams RC, Kendall J, Hamann HA. What happens after distress screening? Patterns of supportive care service utilization among oncology patients identified through a systematic screening protocol. Support Care Cancer. 2016;24(7):2861–8.
- Baker-Glenn EA, Park B, Granger L, Symonds P, Mitchell AJ. Desire for psychological support in cancer patients with depression or distress: validation of a simple help question. Psychooncology. 2011;20(5):525–31.
- Clover KA, Mitchell AJ, Britton B, Carter G. Why do oncology outpatients who report emotional distress decline help? Psychooncology. 2015;24(7): 812–8.
- Dilworth S, Higgins I, Parker V, Kelly B, Turner J. Patient and health professional's perceived barriers to the delivery of psychosocial care to adults with cancer: a systematic review. Psychooncology. 2014;23(6):601–12.
- Schuurhuizen CS, Braamse AM, Beekman AT, Bomhof-Roordink H, Bosmans JE, Cuijpers P, Hoogendoorn AW, Konings IR, van der Linden MH, Neefjes EC, et al. Screening and treatment of psychological distress in patients with metastatic colorectal cancer: study protocol of the TES trial. BMC Cancer. 2015;15:302.
- Singer S, Kuhnt S, Gotze H, Hauss J, Hinz A, Liebmann A, Krauss O, Lehmann A, Schwarz R. Hospital anxiety and depression scale cutoff scores for cancer patients in acute care. Br J Cancer. 2009;100(6):908–12.
- Aaronson NK, Ahmedzai S, Bergman B, Bullinger M, Cull A, Duez NJ, Filiberti A, Flechtner H, Fleishman SB, de Haes JC, et al. The European Organization for Research and Treatment of Cancer QLQ-C30: a quality-of-life instrument for use in international clinical trials in oncology. J Natl Cancer Inst. 1993; 85(5):365–76.
- Schuurhuizen CS, Verheul HM, Braamse AM, Buffart LM, Bloemendal HJ, Dekker J, Konings IR. The predictive value of cumulative toxicity for quality of life in patients with metastatic colorectal cancer during first-line palliative chemotherapy. Cancer Manag Res. 2018;10:3015–21.
- Bouwmans C, De Jong K, Timman R, Zijlstra-Vlasveld M, Van der Feltz-Cornelis C, Tan Swan S, Hakkaart-van Roijen L. Feasibility, reliability and validity of a questionnaire on healthcare consumption and productivity loss in patients with a psychiatric disorder (TiC-P). BMC Health Serv Res. 2013;13:217.
- Zonneveld LN, Sprangers MA, Kooiman CG, van 't Spijker A, Busschbach JJ. Patients with unexplained physical symptoms have poorer quality of life and higher costs than other patient groups: a cross-sectional study on burden. BMC Health Serv Res. 2013;13:520.
- Young J, Badgery-Parker T, Dobbins T, Jorgensen M, Gibbs P, Faragher I, Jones I, Currow D. Comparison of ECOG/WHO performance status and ASA score as a measure of functional status. J Pain Symptom Manag. 2015;49(2): 258–64.
- 26. Stevens JP. Applied multivariate statistics for the social sciences. 4th ed. Mahwah: Lawrence Erlbaum Associates; 2002.
- Vu DH, Muttaqi KM, Agalgaonkar AP. A variance inflation factor and backward elimination based robust regression model for forecasting monthly electricity demand using climatic variables. Appl Energ. 2015;140: 385–94.
- Ellis J, Lin J, Walsh A, Lo C, Shepherd FA, Moore M, Li M, Gagliese L, Zimmermann C, Rodin G. Predictors of referral for specialized psychosocial oncology care in patients with metastatic cancer: the contributions of age, distress, and marital status. J Clin Oncol. 2009;27(5):699–705.
- O'Hea EL, Monahan BR, Cutillo A, Person SD, Grissom G, Boudreaux ED. Predictors of psychological distress and interest in mental health services in individuals with cancer. J Health Psychol. 2016;21(6):1145–56.
- Waller A, Williams A, Groff SL, Bultz BD, Carlson LE. Screening for distress, the sixth vital sign: examining self-referral in people with cancer over a oneyear period. Psychooncology. 2013;22(2):388–95.
- Merckaert I, Libert Y, Messin S, Milani M, Slachmuylder JL, Razavi D. Cancer patients' desire for psychological support: prevalence and implications for screening patients' psychological needs. Psychooncology. 2010;19(2):141–9.
- Kumar P, Casarett D, Corcoran A, Desai K, Li Q, Chen J, Langer C, Mao JJ. Utilization of supportive and palliative care services among oncology outpatients at one academic cancer center: determinants of use and barriers to access. J Palliat Med. 2012;15(8):923–30.

- van Scheppingen C, Schroevers MJ, Smink A, van der Linden YM, Mul VE, Langendijk JA, Coyne JC, Sanderman R. Does screening for distress efficiently uncover meetable unmet needs in cancer patients? Psychooncology. 2011;20(6):655–63.
- Nekolaichuk CL, Cumming C, Turner J, Yushchyshyn A, Sela R. Referral patterns and psychosocial distress in cancer patients accessing a psychooncology counseling service. Psychooncology. 2011;20(3):326–32.
- Herschbach P, Keller M, Knight L, Brandl T, Huber B, Henrich G, Marten-Mittag B. Psychological problems of cancer patients: a cancer distress screening with a cancer-specific questionnaire. Br J Cancer. 2004;91(3):504–11.
- Naughton MJ, Weaver KE. Physical and mental health among cancer survivors: considerations for long-term care and quality of life. N C Med J. 2014;75(4):283–6.
- Steinhauser KE, Christakis NA, Clipp EC, McNeilly M, McIntyre L, Tulsky JA. Factors considered important at the end of life by patients, family, physicians, and other care providers. JAMA. 2000;284(19):2476–82.
- Nolte S, van der Mei SH, Strehl-Schwarz K, Koster J, Bender A, Rose M, Kruse J, Peters EM. Comparison of patient-reported need of psycho-oncologic support and the doctor's perspective: how do they relate to disease severity in melanoma patients? Psychooncology. 2016;25(11):1271–7.
- Hou WK, Lam JH. Resilience in the year after cancer diagnosis: a crosslagged panel analysis of the reciprocity between psychological distress and well-being. J Behav Med. 2014;37(3):391–401.
- Lam WW, Yoon SW, Sze WK, Ng AW, Soong I, Kwong A, Suen D, Tsang J, Yeo W, Wong KY, et al. Comparing the meanings of living with advanced breast cancer between women resilient to distress and women with persistent distress: a qualitative study. Psychooncology. 2017;26(2):255–61.
- Turnbull Macdonald GC, Baldassarre F, Brown P, Hatton-Bauer J, Li M, Green E, Lebel S. Psychosocial care for cancer: a framework to guide practice, and actionable recommendations for Ontario. Curr Oncol. 2012;19(4):209–16.
- Holland JC, Kelly BJ, Weinberger MI. Why psychosocial care is difficult to integrate into routine cancer care: stigma is the elephant in the room. J Natl Compr Cancer Netw. 2010;8(4):362–6.
- Hoekstra-Weebers JEV, M.J.;: Richtlijn 'Detecteren behoefte psychosociale zorg': revisie 2017. Nederlands Tijdschrift voor Oncologie 2017(14):186–192.
- Althubaiti A. Information bias in health research: definition, pitfalls, and adjustment methods. J Multidiscip Health. 2016;9:211–7.
- 45. Dekker J, Beekman AT, Boenink AD, Bomhof-Roordink H, Braamse AM, Collette EH, Huijgens PC, van der Linden MH, van Meijel B, Snoek FJ, et al. Comment on 'psychological distress in patients with cancer: is screening the effective solution?'. Br J Cancer. 2013;108(12):2628–30.
- Abstracts of the 20th World Congress of Psycho-Oncology, 29 October 2 November 2018, Hong Kong, Psychooncology 2018, 27 Suppl 3:3–231.

#### Ready to submit your research? Choose BMC and benefit from:

- fast, convenient online submission
- thorough peer review by experienced researchers in your field
- rapid publication on acceptance
- support for research data, including large and complex data types
- gold Open Access which fosters wider collaboration and increased citations
- maximum visibility for your research: over 100M website views per year

#### At BMC, research is always in progress.

Learn more biomedcentral.com/submissions

