

#### RESEARCH ARTICLE

# Assessment of treatment expectations in women with suspected endometriosis: A psychometric analysis [version 1; peer review: awaiting peer review]

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**V1** First published: 11 Mar 2024, **13**:174

https://doi.org/10.12688/f1000research.145377.1

Latest published: 11 Mar 2024, 13:174

https://doi.org/10.12688/f1000research.145377.1

#### **Abstract**

## **Background**

Treatment expectations influence clinical outcomes in various physical and psychological conditions; however, no studies have explored their role in endometriosis treatment. It is necessary to understand how these expectations can be measured to study treatment expectations and their effects in clinical practice. This study aimed to psychometrically analyze and compare different treatment expectation measurements and describe treatment expectations in women with suspected endometriosis.

#### Method

Analysis of cross-sectional baseline data of a mixed-method clinical observational study of N=699 patients undergoing laparoscopy in Germany. Descriptives, bivariate associations, convergent and discriminant validity of four expectation measurements (Treatment Expectation Questionnaire (TEX-Q); Generic rating scale for previous treatment experiences, treatment expectations, and treatment effects (GEEE); numerical rating scales (NRS) assessing improvement and worsening of endometriosis symptoms, expected Pain Disability Index

#### **Open Peer Review**

Approval Status AWAITING PEER REVIEW

Any reports and responses or comments on the article can be found at the end of the article.

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(PDI); range: 0 to 10) were estimated. A cluster analysis was performed on the three GEEE items.

## **Results**

Most participants expected high improvement (M=6.68 to 7.20, SD =1.90 to 2.09) and low worsening (M=1.09 to 2.52, SD=1.80 to 2.25) of disability from laparoscopy. Participants who expected greater worsening expected more side effects (r=.31 to .60, p<.001). Associations between the positive and negative expectation dimensions, including side effects, were small to non-significant (r =|.24| to .00, p<.001 to.978). Four distinct clusters, described as positive, 'no pain, no gain, 'diminished,' and 'uniform' were found, with a total PVE of 62.2%.

#### **Conclusions**

Women with suspected endometriosis reported positive expectations concerning laparoscopy, but wide ranges indicated interindividual differences. Treatment expectations seem to be a multidimensional construct in this patient group. The investigated measurements did not correlate to the extent that they measured exactly the same construct. The selection of measurements should be carefully considered and adapted for the study purposes. Clusters provide initial indications for individualized interventions that target expectation manipulation.

## **Trial Registration Number**

ID NCT05019612 (ClinicalTrials.gov)

## Keywords

placebo, nocebo, expectancies, laparoscopy, G-EEE, TEX-Q

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Author roles: Meyrose AK: Conceptualization, Data Curation, Formal Analysis, Funding Acquisition, Investigation, Methodology, Project Administration, Validation, Visualization, Writing – Original Draft Preparation; Basedow LA: Conceptualization, Formal Analysis, Methodology, Visualization, Writing – Original Draft Preparation; Hirsing N: Conceptualization, Investigation, Validation, Visualization, Writing – Review & Editing; Buchweitz O: Conceptualization, Investigation, Resources, Supervision, Writing – Review & Editing; Rief W: Conceptualization, Resources, Supervision, Writing – Review & Editing; Nestoriuc Y: Conceptualization, Funding Acquisition, Methodology, Project Administration, Resources, Supervision, Validation, Writing – Review & Editing

**Competing interests:** No competing interests were disclosed.

**Grant information:** This work was funded by internal research funding (IFF2020, 27/05/2020; grant to Prof. Dr. Yvonne Nestoriuc and Dr. Ann-Katrin Meyrose) of the Helmut-Schmidt-University/University of the Federal Armed Forces Hamburg, and was supported by funds from the German Research Foundation (Deutsche Forschungsgemeinschaft, DFG), CRC 289 Treatment Expectation, Project Number 422744262.

The funders had no role in study design, data collection and analysis, decision to publish, or preparation of the manuscript.

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How to cite this article: Meyrose AK, Basedow LA, Hirsing N *et al.* Assessment of treatment expectations in women with suspected endometriosis: A psychometric analysis [version 1; peer review: awaiting peer review] F1000Research 2024, **13**:174 https://doi.org/10.12688/f1000research.145377.1

First published: 11 Mar 2024, 13:174 https://doi.org/10.12688/f1000research.145377.1

#### Introduction

Treatment expectations are important for clinical outcomes in various physical and psychological conditions, <sup>1</sup> such as acute and chronic pain, <sup>2,3</sup> surgeries, <sup>4,5</sup> breast cancer, <sup>6</sup> and psychiatric disorders. <sup>7</sup> Despite the high prevalence of endometriosis (4.4% in the general population<sup>8</sup>) and enormous individual burden, <sup>9</sup> no study has focused on treatment expectations in women with suspected endometriosis. Endometriosis is a chronic inflammatory disease in women of procreative age and is characterized by endometrium-like tissues outside the uterus. <sup>10</sup> The five most prevalent symptoms of endometriosis are dysmenorrhea, abdominal pain, dyspareunia, dyschezia, and dysuria. The German S2k-treatment guideline for endometriosis recommends a laparoscopy, a minimally invasive surgery under general anesthesia, when endocrine therapy has failed. <sup>11</sup> Laparoscopy is associated with short-term symptom improvement, <sup>12</sup> but 20–30% of patients do not respond satisfactorily. <sup>13,14</sup> In summary, current treatment options are inadequate for many patients. <sup>15</sup> A deeper understanding of the role of treatment expectations might help a) clarify why complaints persist and b) develop new intervention avenues to foster positive expectations and prevent nocebo effects. <sup>16</sup>

#### Measurement of expectations

To properly implement and study treatment expectations and their effects in clinical practice, it is necessary to have a good understanding of how these expectations can be measured. This is not as easy as it seems, since treatment expectations can and should be considered multidimensional constructs that differ across specific contexts and time intervals. <sup>17</sup> Specifically, expectations can be positive (e.g., improvement of symptoms) or negative (e.g., worsening of symptoms or side effects), <sup>18</sup> related to treatment processes or outcomes, <sup>19</sup> rated based on valence (more or less important) or probability of occurrence, <sup>20</sup> treatment-specific or related to general symptoms, <sup>21</sup> or different in terms of being real, that is, plausible or ideal expectations. <sup>17,19</sup> This complexity of the 'treatment expectation' construct leads to pronounced heterogeneity in available instruments. <sup>22–24</sup> While single-item measures have been the most popular historically, <sup>20,23,25</sup> recent advances have led to the development of multidimensional treatment-expectation questionnaires. <sup>22,26</sup> Although this development circumvents several issues in the assessment of treatment expectations, <sup>17</sup> an ideal method of expectation measurement has not been developed. Several issues remain understudied and are fruitful targets for continued psychometric research, such as the contrast between context specific and general expectations.

## Aims and hypotheses

The main aim of this study was to psychometrically analyze and compare different treatment expectation measurements in a large sample of *N*=699 women with suspected endometriosis. Additionally, the study aimed to describe and exploratively cluster treatment expectations in this burdened and understudied patient group. We hypothesise that:

- i) Treatment expectations are a multidimensional construct; that is, intercorrelations of different scales and dimensions of the *Treatment Expectation Questionnaire* (TEX-Q)<sup>22</sup> and the three expectation items of the *Generic rating scale for previous treatment experiences, treatment expectations, and treatment effects* (GEEE<sup>26</sup>) will be heterogeneous.
- ii) Treatment expectation measurements correlate higher with each other in terms of convergent validity compared to other psychological measurements assessed (i.e., disability, severity of symptoms, depressive symptoms, anxiety, and catastrophizing pain) in terms of discriminant validity.

## Methods

This study used data from the baseline assessment of a mixed-method clinical observational study<sup>27</sup> that focused on patients undergoing laparoscopy for suspected endometriosis. Women with endometriosis reported their endometriosis-related disability, complaints, and a priori specified predictors once before and eight times after laparoscopy, covering trajectories over a 12-month period. This observational study was registered at ClinicalTrials.gov (ID NCT05019612), but baseline analyses were not described.

## **Participants**

The target population included adult women with endometriosis-related complaints. Further inclusion criteria were sufficient German language skills, female sex, informed consent for study participation, and indications for laparoscopy. For these analyses, N=699 patients were included, irrespective of the actual surgery or clinical diagnosis after laparoscopy.

## Recruitment and procedure

Women with an appointment for laparoscopy in a specialized center for surgical endoscopy and endometriosis (Frauenklinik an der Elbe, Germany) were informed about the study by phone. If interested, the women were referred to the baseline online survey. The online survey included written information about the study and checked the inclusion

criteria. Written informed consent for the online survey, storage and processing of data was obtained online from all participants, and baseline assessment was initiated. An individualized study code was used to store the data. Patients completed the survey between 25<sup>th</sup> of August 2021 to 27<sup>th</sup> of June 2023 until the required sample size of longitudinally participating patients with diagnosed endometriosis was reached. Interested patients were sent email reminders to encourage their participation. Detailed information on the study design, postoperative assessments, and further efforts to address potential bias are described in the study protocol.<sup>27</sup>

The authors assert that all procedures contributing to this work comply with the ethical standards of the relevant national and institutional committees on human experimentation and the Helsinki Declaration of 1975, as revised in 2008. All procedures involving patients were approved by the Psychotherapeutenkammer Hamburg, Germany (ROXWELL-2021-HH, 25<sup>th</sup> of June 2021).

#### Measurements

Treatment expectations about laparoscopy were assessed by four self-reported measurements (Treatment Expectation Questionnaire; three items assessing treatment expectations of the Generic rating scale for previous treatment experiences, treatment expectations, and treatment effects; self-constructed numerical rating scales; and expected Pain Disability Index). The Treatment Expectation Questionnaire (TEX-Q)<sup>22,28</sup> comprises 15 items of six dimensions with 11 response options each. The mean score ranged from 0 to 10. Higher scores indicated more positive treatment expectations for the total score and dimensions of 'treatment benefit', 'positive impact', 'process', and 'behavioral control'. Higher scores indicate more negative treatment expectations for the 'adverse events' and 'negative impact' dimensions. In this study, Cronbach's  $\alpha$  of the total score was.82 and ranged between.72 and.91 for the dimensions in accordance with the validation sample.<sup>28</sup>

Three items assessing treatment expectations of the Generic rating scale for previous treatment experiences, treatment expectations, and treatment effects (GEEE)<sup>26</sup> were used, focusing on expected improvement of disability, worsening of disability, and side effects from laparoscopy with 11 response options (0=no improvement/impairment to 10=greatest improvement/worsening imaginable; 0=no complaints to 10=greatest complaints imaginable). No sum or mean scores were obtained.

Ten self-constructed numerical rating scales (NRS) with 11 response options were used to assess the expected improvement and worsening of the five most prevalent endometriosis symptoms (dysmenorrhea, pelvic/abdominal pain, dyspareunia, dyschezia, and dysuria). The wording of items and response options (0=no improvement/impairment to 10=greatest improvement/worsening imaginable) were derived from the GEEE in accordance with the recommendations for the assessment of pain. No sum or mean scores were obtained.

Expected endometriosis-related pain disability was assessed using the German version of the Pain Disability Index $^{29,30}$  adapted to expected disabilities after laparoscopy (expected PDI). The expected PDI covers seven items with 11 response options (0=no disability to 10=total disability). The sum (range: 0-70) and mean scores (0-10) were calculated. Higher scores indicate higher expected disability. The mean scores were mainly used because of their better comparability to the other expectation measurements in this study. The Cronbach's  $\alpha$  was.93 in this study. The expected change in disability by laparoscopy was defined as the difference between endometriosis-related disability before laparoscopy and expected endometriosis-related disability after laparoscopy ( $\Delta$ PDI score=PDI – expected PDI; theoretically ranging from -70 to 70 for the sum and -10 to 10 for the mean score). Positive scores indicate a positive change in disability by laparoscopy, that is, improvement, whereas negative scores indicate a negative change in disability, that is, worsening.

Further, self-reported psychological measurements were used to describe the sample and estimate discriminant validity. Endometriosis-related disability was assessed using the PDI $^{29,30}$  covering seven items with 11 response options (0=no to 10=total disability). The introductory text was adapted to *endometriosis-related* disability. The sum (range: 0-70) and mean scores (0-10) were calculated. Higher scores indicate higher disability. Cronbach's  $\alpha$  was .85 in this study. Disability will be compared to women of the German general population.<sup>31</sup>

The severity of endometriosis-related symptoms was assessed by five NRS referring to the five most prevalent endometriosis symptoms (dysmenorrhea, pelvic/abdominal pain, dyspareunia, dyschezia, and dysuria) with 11 response options (0=no pain to 10=worst pain imaginable). To summarize symptoms, the maximum severity of symptoms score was calculated using the stated maximum severity of symptoms within the five NRS, for example, if someone selected response options between 2 and 8 for the five endometriosis symptoms, the maximum severity of symptoms score was 8.

Depressive symptoms and anxiety were assessed using the Patient Health Questionnaire (PHQ-4)<sup>32</sup> which covers four items with four response options (0=not at all, 1=several days, 2=more than half the days, 3=nearly every day). Sum scores for the depressive symptoms and anxiety subscales ranged from 0 to 6. Higher scores indicate more depressive symptoms and anxiety, respectively. Cronbach's  $\alpha$  was .77 for depressive symptoms and .76 for anxiety in this study. A cutoff score of 3 or greater indicates good sensitivity and specificity for detecting major depression<sup>33</sup> and anxiety disorders.<sup>34</sup>

Pain catastrophizing was assessed by the subscale catastrophizing of the Coping Strategies Questionnaire  $(CSQ)^{35}$  which covers six items with seven response options each (0=never do that to 6=always do that). The sum score ranges from 0 to 36, with higher scores indicating a higher level of catastrophizing pain. The Cronbach's  $\alpha$  was .88 in this study.

#### Statistical analysis

Frequencies and descriptive statistics (mean, standard deviation, median, range, skewness, and kurtosis) were calculated to describe the sample. Missing data are also reported. Distributions of treatment expectation measurements are displayed by raincloud plots using the R packages ggplot2, ggdist, ggforce, gghalves, haven, RcolorBrewer. Bivariate associations were estimated using Pearson correlation coefficients and displayed as a correlation matrix heatmap using the R package ggcorrplot. Convergent validity was determined by bivariate associations of all treatment expectation measurements, discriminant validity by bivariate associations of treatment expectation measurements, and psychological constructs. According to Cohen, Pearson's r=.1-.3 is interpreted as small, r=.3-.5 as small, and small and small are small and small by small and small are small and small and small are small and small are small and small are small and small and small are small and small are small and small and small are small and small are small and small and small are small and small and small are small and small and small and small are small and small and small and small are small and small and small and small and small are small and small and small are small and small and small and small and small are small and small and small and small and small and small and small are small and small are small and small and small and small and small and small and small ana

#### Results

#### Descriptive characteristics

Overall, n=2,361 people clicked the survey link, n=1,145 began the online survey, n=1,098 met the inclusion criteria, n=1,082 gave informed consent, and n=699 completed the online survey. More details on the flow of participants are displayed in Figure 1.

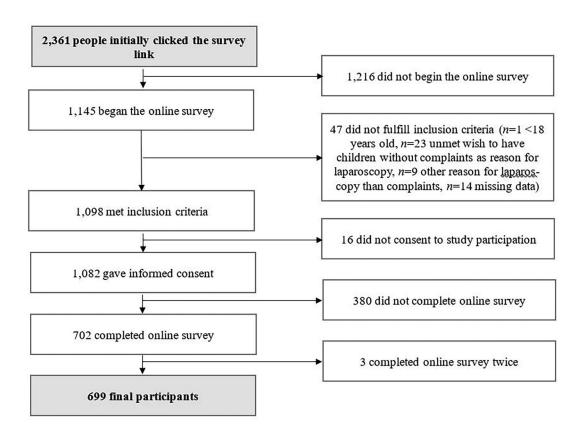


Figure 1. Flow chart of study participation.

Table 1. Descriptive characteristics of the sample under analysis (N=699).

	(%) u	×	as	Ма	Observed range	Potential range	Skew	Kurt
Gender								
Female	686 (98.1)							
Non-binary	13 (1.9)							
Age in years	669	29.90	6.45	29.50	18-51	>18	0.47	-0.21
Migrant background								
Yes	129 (22.5)							
No	445 (77.5)							
Educational level								
No school degree	2 (0.4)							
In school	8 (1.1)							
Lower secondary school	13 (1.9)							
Secondary school	170 (24.3)							
Higher entrance certificate	274 (39.2)							
University degree	232 (33.2)							
Reason for laparoscopy								
Complaints	549 (77.5)							
Complaints with an unmet wish to have children	150 (21.5)							
Able to work despite endometriosis-related complaints								
Yes	623 (89.1)							
No	76 (10.9)							
Severity of symptoms (NRS)								
Dysmenorrhoea <sup>a</sup>	624 (89.3)	6.85	1.95	7.00	0-10	0-10	-0.80	0.51
Abdomen	669	5.72	2.27	00.9	0-10	0-10	-0.50	-0.32
Sexual intercourse <sup>b</sup>	476 (68.1)	3.61	2.72	3.50	0-10	0-10	0.31	-0.85
Dysuria	669	1.43	2.10	0.00	0-10	0-10	1.50	1.48
Dyschezia	669	2.52	2.60	2.00	0-10	0-10	0.76	-0.44
Max. severity of symptoms	669	7.18	1.81	8.00	1-10	0-10	-0.82	0.55
Disability (PDI)								
Total sum score	669	30.52	13.66	31.00	0-70	0-70	0.01	-0.31
Total mean score	669	4.36	1.95	4.43	0-10	0-10	0.01	-0.31
Depressive symptoms (PHQ-4)	669	2.56	1.40	2.00	9-0	9-0	0.53	-0.03
Anxiety (PHQ-4)	669	2.44	1.63	2.00	9-0	9-0	0.43	-0.53

Table 1. Continued

	n (%)	N	SD	Мд	Observed range	Potential range	Skew	Kurt
Catastrophizing pain (CSQ)	669	18.26	6.91	18.00	6-35	0.36	0.25	-0.62
TREATMENT EXPECTATIONS								
TEX-Q								
Total score	669	6.73	1.28	6.73	2.40-10	0-10	-0.19	0.03
Treatment benefit	669	7.15	1.90	7.33	0-10	0-10	-0.67	0.20
Positive impact	669	89.9	2.35	7.00	0-10	0-10	-0.64	-0.17
Adverse events	669	3.67	1.93	3.33	29.6-0	0-10	0.34	-0.34
Negative impact	669	2.52	2.25	2.00	0-10	0-10	08.0	-0.08
Process	669	68.9	1.92	7.00	0-10	0-10	-0.30	-0.32
Behavioral control	669	5.87	2.44	00.9	0-10	0-10	-0.33	-0.52
Generic rating scales (GEEE)								
Improvement	669	7.20	2.09	8.00	0-10	0-10	-0.84	0.67
Worsening	669	1.09	1.79	0.00	0-10	0-10	2.17	5.21
Side effects	669	3.72	2.33	3.00	0-10	0-10	0:30	-0.62
Expected severity of symptoms (NRS)								
Dysmenorrhoea improvement <sup>a</sup>	624 (89.3)	6.70	2.34	7.00	0-10	0-10	-0.73	0.19
Dysmenorrhoea worsening <sup>a</sup>	624 (89.3)	0.0.92	1.79	00:00	0-10	0-10	2.68	8.10
Abdomen improvement	669	6.94	2.35	7.00	0-10	0-10	-0.86	0.51
Abdomen worsening	669	1.05	2.01	00:00	0-10	0-10	2.50	6.21
Sexual intercourse improvement <sup>b</sup>	476 (68.1)	5.58	3.64	7.00	0-10	0-10	-0.38	-1.29
Sexual intercourse worsening <sup>b</sup>	476 (68.1)	0.91	2.04	00:00	0-10	0-10	2.79	7.60
Dysuria improvement	669	3.73	4.04	2.00	0-10	0-10	0.41	-1.53
Dysuria worsening	669	09:0	1.62	0.00	0-10	0-10	3.45	12.58
Dyschezia improvement	669	4.67	3.87	2.00	0-10	0-10	00.00	-1.58
Dyschezia worsening	669	0.70	1.78	0.00	0-10	0-10	3.17	10.32
Expected disability (expected PDI)								
Total sum score	669	15.41	13.77	12.00	99-0	0-70	1.07	0.52
∆Total sum score <sup>c</sup>	669	15.10	14.12	14.00	-36-65	-70-70	0.16	0.34
Total mean score	669	2.20	1.97	1.71	0-9.43	0-10	1.07	0.52
∆Total mean score <sup>c</sup>	669	2.16	2.02	2.00	-5.14-9.29	-10-10	0.16	0.34

Note. M, mean; 5D, standard deviation: Md, median; Skew, skewness; Kurt, kurtosis; PDI, Pain Disability Index; NRS, numerical rating scale; PHQ-4, Patient Health Questionnaire, 4-item version; CSQ, Coping Strategies Questionnaire; GEEE, Generic Rating Scale for Treatment Expectations; TEX-Q, Treatment Expectation Questionnaire; ∆, difference score.

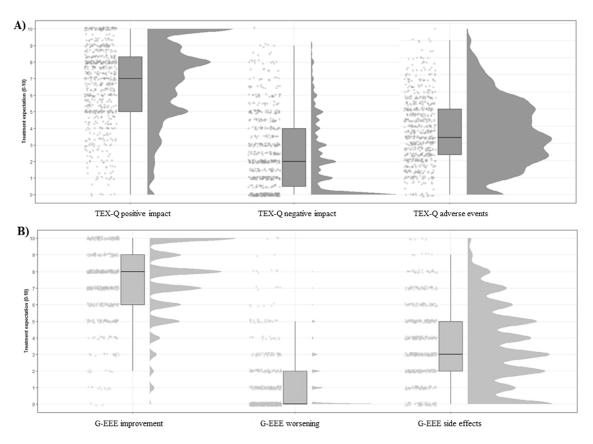
<sup>a</sup>Menstrual bleeding was not applicable for n=75.

<sup>b</sup>Sexual intercourse was not applicable for n=223.

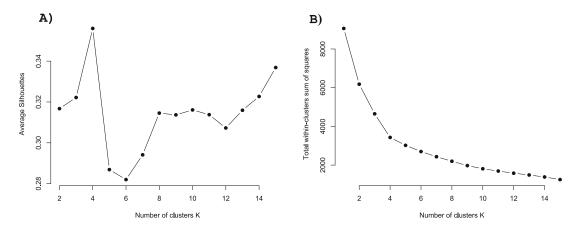
<sup>c</sup>n=74 (10.6%) had negative values, that is, expected worsening of disability; n=22 (3.1%) had a value of zero, that is, expected no change in disability.

The ages of the 699 participants with suspected endometriosis ranged from 18 to 51 years (M=29.90, SD=6.45). About one-fifth had a migrant background, i.e. participants or one or both parents were not born in Germany, and 39.2% had acquired a higher entrance certificate (German 'Abitur') and 33.2% had a university degree. The reason for laparoscopy was endometriosis-related complaints with (21.5%) or without (77.5%) an unmet wish to have children. Overall, participants were enormously burdened by endometriosis-related symptoms before laparoscopy (PDI sum score: M=30.52, SD=13.66) compared with women in the general population in Germany (M=6.9, SD=11.1, N=1,368 $^{31}$ ; d=3.14). On average, the maximum severity within the five endometriosis-related symptom scales (NRS) was M=7.18 (SD=1.81, potential range: 0-10). More than half (54.8%) showed signs of major depression (41.6%, M=2.56, SD=1.40) and/or anxiety disorder (42.6%, M=2.44, SD=1.63) according to the PHQ-4. Table 1 presents more information on the characteristics of the sample under analysis and the descriptive statistics of all the treatment expectation measurements.

In general, women with suspected endometriosis had rather positive treatment expectations concerning laparoscopy. However, the wide ranges of all measurements, mostly covering the total potential range, indicate pronounced interindividual differences. Positive dimensions (TEX-Q; treatment benefit, positive impact) and items (GEEE, NRS), such as expected improvement, were rated with higher values (M=6.68 to 7.20, SD=1.90 to 2.35) and were distributed slightly left-skewed (skewness: -0.84 to -0.64). Negative dimensions (TEX-Q; adverse events, negative impact) and items (GEEE, NRS) such as expected worsening (M=0.92 to 1.09, SD=1.79) or side effects (M=3.67 and 3.72, SD=1.93 and 2.33) were rated with lower values and distributed clearly right-skewed (0.30 to 3.45). Expected disability (PDI total mean score) after laparoscopy ranged from 0 to 9.43 (M=2.20, SD=1.97) and was right-skewed (1.07). Expected change of disability ( $\Delta$ total mean score) ranged from -5.14 (medium worsening) to 9.29 (high improvement). Scores tended to be normally distributed based on graphical examination and skewness and kurtosis parameters, whereas the Kolmogorov-Smirnov test was significant (D(699)=.05, p<0.001). The distributions of some exemplary expectation measurements are shown in Figure 2.



**Figure 2. Distributions of treatment expectation measurements using raincloud plots.** *Note.* A: TEX-Q subscales: positive impact, negative impact, and adverse events. B: GEEE scale improvement, worsening, and side effects.



**Figure 3. Silhouette and elbow plots for the exploration of possible factors.** *Note.* A: Silhouette plot showing the average silhouette score in relation to the number of clusters. B: Elbow plot showing the total within-cluster sum of squares for the number of potential clusters.

## Cluster of treatment expectations

Figure 3 shows the resulting elbow and silhouette plots for potential clustering of the three GEEE items (symptom improvement, symptom worsening, and side effects). Both plots suggest k=4 is the optimal number of clusters. K-means clustering with k=4 resulted in a proportion of explained variance of 62.2%. The details of these clusters are listed in Table 2. Descriptively, the clusters can be described as 'positive' (high improvement expectation), 'no pain, no gain' (high improvement and side effect expectation), 'diminished' (rather low expectations on every dimension), 'uniform' (equally high expectations in every direction) (see Figure 4).

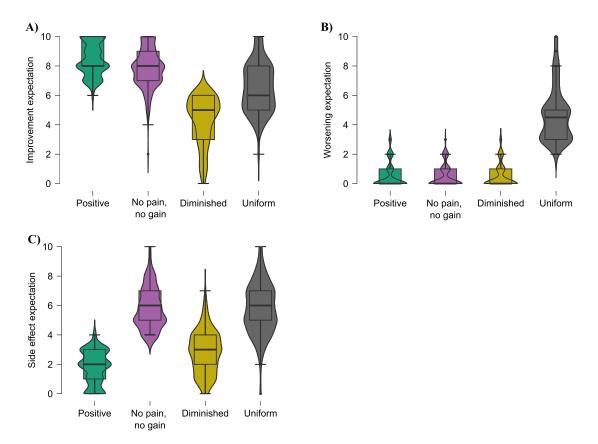
Table 2. Characteristics of the four identified clusters.

	Cluster 'positive' (n=260)	Cluster 'no pain, no gain' ( <i>n</i> =204)	Cluster 'diminished' (n=137)	Cluster 'uniform' (n=98)
Cluster centroids				
Generic rating scales (GEEE) M (SD)				
Improvement	8.46 (1.14)	7.85 (1.35)	4.30 (1.67)	6.55 (1.67)
Worsening	0.53 (0.88)	0.47 (0.75)	0.48 (0.81)	4.71 (1.87)
Side effects	1.76 (1.15)	5.84 (1.41)	2.82 (1.45)	5.77 (1.75)
Demographics per cluster				
Mean age in years (SD)	29.9 (6.6)	30.5 (6.6)	29.2 (6.5)	29.6 (5.7)
<b>N</b> female (divers)	255 (5)	202 (2)	132 (5)	97 (1)
<b>N</b> unable to work because of endometriosis-related complaints (%)	21 (8.1)	16 (7.8)	16 (11.7)	23 (23.5)
Migrant background N (%)				
Yes	53 (25.5)	29 (16.8)	19 (17.0)	28 (34.6)
No	155 (74.5)	144 (83.2)	93 (83.0)	53 (65.4)
Educational level N (%)				
No school degree	2 (0.8)	0	0	0
In school	3 (1.2)	3 (1.5)	1 (0.7)	1 (1.0)
Lower secondary school	10 (3.8)	1 (0.5)	2 (1.5)	0
Secondary school	73 (28.1)	48 (23.5)	16 (11.7)	33 (33.7)
Higher entrance certificate	88 (33.9)	79 (38.7)	63 (46.0)	44 (34.9)
University degree	84 (32.3)	73 (35.8)	55 (40.1)	20 (20.4)

Table 2. Continued

	Cluster 'positive' (n=260)	Cluster 'no pain, no gain' ( <i>n</i> =204)	Cluster 'diminished' (n=137)	Cluster 'uniform' (n=98)
Reason for Laparoscopy $N\left(\%\right)$				
Complaints	201 (77.3)	156 (76.5)	111 (81.0)	81 (82.7)
Complaints with an unmet wish to have children	59 (22.7)	48 (23.5)	26 (19.0)	17 (17.3)
Previous experience with laparoscop	y N (%)			
Yes, within last 12 months	8 (3.1)	6 (2.9)	3 (2.2)	6 (6.1)
Yes, more than 12 months ago	55 (21.2)	32 (15.7)	18 (13.1)	21 (21.4)
No	197 (75.8)	166 (81.4)	116 (84.7)	71 (72.4)
Disability (PDI)				
Total mean score (SD)	4.3 (1.8)	4.3 (2.0)	4.0 (2.0)	5.1 (2.0)
Depressive symptoms (PHQ-4)				
Total mean score (SD)	2.5 (1.4)	2.5 (1.3)	2.5 (1.5)	2.9 (1.4)
Anxiety (PHQ-4)				
Total mean score (SD)	2.4 (1.7)	2.2 (1.5)	2.5 (1.8)	2.9 (1.6)
Catastrophizing pain (CSQ)				
Total mean score (SD)	17.3 (7.6)	17.1 (7.7)	16.2 (8.2)	19.8 (7.7)

Note. SD, standard deviation; GEEE, Generic Rating Scale for Treatment Expectations; PDI, Pain Disability Index; NRS, numerical rating scale; PHQ-4, Patient Health Questionnaire, 4-item version; CSQ, Coping Strategies Questionnaire.



**Figure 4. Distribution of the GEEE values for each cluster.** *Note.* A: Average GEEE improvement expectation scores for each of the four clusters. B: Average GEEE worsening expectation scores for each of the four clusters. C: Average GEEE side effect expectation scores for each of the four clusters.

## Convergent validity: Bivariate associations of all treatment expectation measurements

All associations of the treatment expectation measurements are presented in Figure 5, with the underlying statistics in Table 3. The dimensions of 'treatment benefit' and 'positive impact' of the TEX-Q were largely associated with each other (r=.75, p<.001), the total score (r=.76, p<.001 and r=.73, p<.001), and with a medium effect size with the expected process (r=.49, p<.001 and r=.42, p<.001, respectively). Negative dimensions ('adverse events' and 'negative impact') were largely associated with each other (r=.60, p<.001) and the total score (r=-.51 to r=-40, p<.001), but correlations with positive dimensions were non-significant to small. The correlations of the GEEE items were non-significant to small. Participants who expected more worsening from laparoscopy also expected more side effects (r=.31, p<.001) and less improvement (r=-.09, p=.016), but the effect sizes were small. The expected improvement and side effects of laparoscopy were not associated (r=-.07, p=.061). Correlations of expected worsening and improvement of each endometriosis-related symptom (NRS) were non-significant to small (r=.15 to .24, p<.001), whereas some expected changes in specific endometriosis-related symptoms were highly associated with the same expected change of other symptoms, for example, improvement of dysmenorrhea and abdominal pain, worsening of dysuria, and dyschezia (r=.69 to.73, p<.001). The correlation between expected disability (expected PDI) and change in disability ( $\Delta$ total score) was high (r=-.52, p<.001); participants who expected more disability after laparoscopy also expected worsening or a smaller reduction in disability.

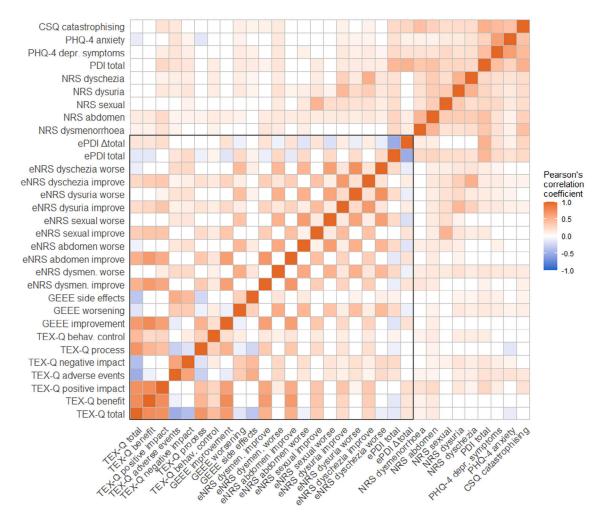


Figure 5. Heatmap representing pearson correlation coefficients between treatment expectation measurements and further psychological constructs. *Note.* Red indicates positive correlations, and blue indicates negative correlations. Non-significant correlations are white. The box highlights the correlations between the treatment expectation measurements (convergent validity). More information on each measurement is provided in the Methods section *Measurements*. The underlying statistics are presented in Table 3. TEX-Q, Treatment Expectation Questionnaire; GEEE, Generic Rating Scale for Treatment Expectations; eNRS, expected numerical rating scales; ePDI, expected Pain Disability Index;  $\Delta$ , difference score; NRS, numerical rating scales; PDI, Pain Disability Index; PHQ-4, Patient Health Questionnaire, 4-item version; CSQ, Coping Strategies Questionnaire.

Table 3. Correlation matrix including all treatment expectation measurements and further psychological constructs under analysis (N=699).

	31.																						
	30.																						
	.62																						
	78.																						
	27.																						
	. 56.																						
	25.																						
	24.																						
	23.																						
	72.																						-
	21.																					-	,520 ,001
	20.																				-	.237 001	-115 -002
	6.																			-	.183 001	.007	.002
	8.																		-	.206 001	.732 4.001	.231 4.001	095
	17.																	-	.244 <.001	.001 001	.001 001	.405	.115 .002
	16.																-	.001	.605 <.001	.009	.628 001	.001 001	
	15.															-	.001	.442 4.001	.087	.399 <.001	.076 .097	052	 181. 100.
	4														-	.031	.647	.00 .00	.522 <.001	.066	.495 <.001	.236 4.001	162 001
	13.													-	.014	.418 001	900-	,294 ,001	210.	.321 <.001	945	·.126 <.001	 .00.
	12.												-	.043	.575 001	.054	.509 .001	.157 001	.519 001	.166 001	.470 001	.249 <.001	083 .038
	Ę											-	.065	.691 001	.597	.324 <.001	.690	.295 001	.056	.316 <.001	.051	095 710.	.4.00.
	-01										-	.942	.167	016	.137	002	.081	.070	.094 0.13	.029	.004 .004	.093	.040
	ъ									-	309 .001	.013	.544 001	033	.465 001	.030	393 .	.001	.435 001	.00.	.452 .001	.238 .001	088
	∞i								-	091 016	071	.626 <.001	.011	.643 .001	030	.365 <.001	006 .904	.255 <.001	.565	.290 4.001	000.	161 0001	.00. 195
	7.							-	.205 .001	.118	.014	145 <.001	.095 .018		.045	.002	.294	.001	.514	.001 2.001	.060	.076 .044	.320
	9						-	308 001	.507 001	092 .015	-,239	.352 <.001	025	.369 001	.034	.243 <.001	054 .239	.182 <.001	003	.206 .001	032 .392	087 .022	.070
	ιń					-	168 001	.002	043 .255	.340 4.001	.449 .001	.181	.251	016	.207 <.001	.042	.174 001	.003	.198	.003	.254 <.001	.245 <.001	073
	4				-	601	288	.113	109	.338 <.001	.544 .001	.016	.246 <.001	043	.163 <.001	.013	.157 4.001	600.	.180 001	.079 7E0.	.245 <.001	.172 2.001	.016
	mi			-	.001	.070	.419 4.001	.284 4.001	.625 4.001	.063	054	.586 4.001	.135 001	.549 <.001	.231	.381 4.001	.053	.381 4.001	.032	362 <.001	.086	.468	.001 001
	2		-	.748 <.001	077 .042	022	.491 <.001	.243 <.001	.001 	057	068	.663	.001	.666 001	016 .670	383	.013	.270 001	.883	.327 <.001	940	.142 <.001	.173
пp	÷	-	.762 <.001	.731 <.001	509	401	.706 001	.448 .001	.007 001	-,164 -,001	.354 <.001	.505 001	064	.534 <.001	081	.332 <.001	063	.232 <.001	0.64	.259 <.001	095 .012	.140 .001	.169 .000
	Measurement	1. TEX-Q Total score	2. TEX-Q Treatment benefit	3. TEX-Q Positive impact	4. TEX-Q Adverse events	5. TEX-Q Negative Impact	6. TEX-Q Process	7. TEX-Q Behavioral control	8. GEEE Improvement	9. GEEE Worsening	10. GEEE Side effects	11. eNRS Dysmenorrhoea improvement	12. eNRS Dysmenorrhoea worsening	13. eNRS Abdomen improvement	14. eNRS Abdomen worsening	15. eNRS Sexual intercourse improvement	16. eNRS Sexual intercourse worsening	17. eNRS Dysuria improvement	18. eNRS Dysuria worsening	19. eNRS Dyschezia improvement	20. eNRS Dyschezia worsening	21. ePDI Total mean score	22. ePDI ∆Total mean score

Table 3. Continued

	31.									-
	30.								-	.402 <.001
	29.							-	.637 <.001	.454 <.001
	28.						-	.435 <.001	.290 <.001	.520 <.001
	27.					-	.255 <.001	.210 <.001	.181 <.001	.227 <.001
	.92				-	.507 <.001	.271 <.001	.194 <.001	.136 <.001	.232 <.001
	25.			-	.343 <.001	.219 <.001	.258 <.001	.252 <.001	.002	.219 <.001
	24.		-	.335 <.001	.332 <.001	.334 <.001	.409	.178 001	.093	.297 <.001
	23.	-	.482 <.001	.180 <.001	.202 <.001	.221 <.001	.374 <.001	.187 <.001	10. 10.	.417 <.001
	22.	.003	.001	.504	.050	.062	.509 <.001	.190 001	600.	.273 <.001
	21.	.248 <.001	.283 <.001	.001 001	.218 <.001	.00. 001	.471 <.001	.237 001	.186 001	.236 <.001
	20.	069	.005	.007	.160 001	.001 001	121.	.050	.041	.051
	6.	.060	.160	.191 001	.319 <.001	.527 <.001	.130	.133	.085	.092
	8.	.056	.002	.185 001	.245 <.001	.143 <.001	.134 001	.096	.039	.067
	17.	.071	.155 001	.227 001	.473 <.001	.297 <.001	.151 -4.001	.153 <.001	.006	.094
	16.	.021	.002	.217 <.001	.035	.079	.395	.032	.018	.032
	5.	.637	.004	.502 <.001	171.	.010 010	.002	.084	.987	.104
	4	.110 006	.145 <.001	.001	.005	.034	.070	.008 008	.047	690.
	<del>.</del>	.051	.131 <.001	.071	.026	.345	720.	.801	.063 .096	.030
	12.	.133 <.001	.160 001	.003	.194 001	.132 <.001	.168 000	.158 000	.152 <.001	.143 <.001
	Ξ.	.171. 2.001	.001	004	.073 070.	.068	.024	.428	026	.103 010
	-01	.045	.133	.057	.084	.074	.135 <.001	600.	.064 94	.076 .045
	ക്	.064	.089	.092	.145 <.001	.106	.149 <.001	.128 .001	.091	.003
	∞i	.061	.003	.036	.067	.042	.039	.006	061	.052
	7.	.028	.134 <.001	.113	.001	.124 001	.116	.005	018	.050
	9	.856	.094	.038	.030	.010	015	056	-, 153 -,001	-019
	ιή	.031	.090	.093	.174 001	.119	171.	.005	.081	.067 770.
	4	.084	.042	.139	.00. 121	.003	.190 -100.	.001 001	.178 001	.161 00.
	mi	.191 001	.236 <.001	.080	.161 001	.086	.257 001	.002	.548	.193 00.
	7	.090	.140	.031	-035	.568	.035	.748	058	.193
цb		.088	.137	.024	.030	.010	.034	041 .274	-117 -002	.030
	Measurement	23. NRS Dysmenorrhoea	24. NRS Abdomen	25. NRS Sexual intercourse	26. NRS Dysuria	27. NRS Dyschezia	28. PDI total mean score	29. Depressive symptoms (PHQ-4)	30. Anxiety (PHQ-4)	31. Catastrophizing pain (CSQ)

Note. The significant correlations are indicated in bold. TEX-Q, Treatment Expectation Questionnaire; GEEE, Generic Rating Scale for Treatment Expectations; eNRS, expected numerical rating scales; ePDI, Pain Disability Index; PHQ-4, Patient Health Questionnaire, 4-item version; CSQ, Coping Strategies Questionnaire.

The positive dimensions of treatment expectations regarding laparoscopy, the expected process, and the total score of the TEX-Q were highly and positively correlated with the expected improvement in disability as measured by the GEEE (r=.51 to .77, p<.001). Medium-sized positive correlations were found between the negative dimensions of the TEX-Q and expected worsening (r=.34, p<.001 and r=.34, p<.001) and expected side effects (r=.45 to.55, p<.001), as measured by the GEEE.

Considering the expected improvement and worsening of the severity of endometriosis-related symptoms (NRS), participants who expected more treatment benefit, positive impact from the laparoscopy, and had more positive overall treatment expectations measured by the TEX-Q, also reported more improvement in the severity of symptoms, especially dysmenorrhea (r=.51 to .66, p<.001) and abdominal pain (r=.53 to .67, p<.001). Negative dimensions of the TEX-Q were significantly correlated with the expected worsening of symptom severity, but the effect sizes were small (r=.16 to .25, p<.001). Additionally, the expected improvement and worsening of disability measured by the GEEE were positively correlated with the expected improvement and worsening of the severity of endometriosis-related symptoms (NRS), respectively, ranging from rather high correlations for dysmenorrhea (improvement: r=.63 and worsening: r=.54, p<.001) and abdominal pain (r=.64 and .47, p<.001) to rather medium correlations for pain during sexual intercourse (r=.37 and .39, p<.001), dysuria (r=.26 and .44, p<.001), and dyschezia (r=.29 and .45, p<.001).

Considering the expected disability after laparoscopy measured using the expected PDI, all correlations with other treatment expectation measurements were small. Participants who expected lower disability after laparoscopy had more positive treatment expectations overall (r=-.14, p<-.001) and expected fewer adverse events and negative impact measured by the TEX-Q (r=.17 and.26, p<-.001). Participants who expected a higher positive change in disability ( $\Delta$ total score) by laparoscopy (i.e., improvement of disability) also expected more benefit from treatment, more positive impact, and more positive treatment expectations overall (r=.17 to.22, p<-.001). No associations were found with the other TEX-Q dimensions. Additionally, the correlations between the expected improvement and worsening of disability measured by the GEEE and expected disability after laparoscopy (r=-.16, p<-.001 and r=-.24, p<-.001) and expected changes in disability (r=.20, p<-.001 and r=-.09, p=.02) measured by the PDI were small, although both measurements explicitly referred to endometriosis-related disability. Similarly, almost the same associations regarding direction and effect size were found between expected disability and change in disability measured by the expected PDI and expected improvement and worsening of almost all endometriosis-related symptoms (NRS).

## Discriminant validity: Bivariate associations of treatment expectation measurements and further psychological constructs

Overall, correlations between all treatment expectation measurements and investigated further psychological constructs, that is, preoperative disability, severity of the five most prevalent endometriosis-related symptoms, depressive symptoms, anxiety, and catastrophizing pain, were non-significant to small, with a few exceptions of medium-sized associations. Correlations between treatment expectations measured by the TEX-Q and all psychological constructs were non-significant to small. The most notable correlations were as follows: participants who experienced more preoperative disability and a higher severity of abdominal pain expected a higher positive impact from laparoscopy (r=.26 and .24, p<.001). Participants who reported more depressive symptoms and anxiety expected more adverse events (r=.18 and .18, p<.001).

The expected improvement, worsening, and side effects measured by the GEEE were not correlated with any of the psychological constructs to a relevant extent (r<.13). Correlations between expected improvement and worsening of the severity of the five most prevalent endometriosis-related symptoms and all further psychological constructs were non-significant to small, despite a few exceptions. Participants who experienced higher pain during sexual intercourse, more severe dysuria, and more severe dyschezia expected considerably more improvement in these specific complaints (r=.47 to .53, p<.001). Expected disability after laparoscopy and expected change in disability measured by PDI were highly correlated with preoperative disability. Participants who experienced more preoperative disability expected a higher disability after laparoscopy (r=.47, p<.001) and a higher change, that is, improvement of disability by laparoscopy (r=.51, p<.001). Additionally, participants who reported more severe endometriosis-related symptoms, more depressive symptoms, and more anxiety expected a higher disability after laparoscopy (r=.19 to .29, p<.001). Participants with a stronger tendency to catastrophize pain expected both a higher disability after laparoscopy and a higher chance, that is, improvement of disability from laparoscopy (r=.24 and .27, p<.001).

#### Discussion

#### Summary

Women with suspected endometriosis reported rather positive treatment expectations concerning laparoscopy, but wide ranges emphasize differences between participants. The negative dimensions indicate major floor effects. As hypothesized, low correlations between the dimensions of treatment expectation measurements indicate that treatment

expectation also seems to be a multidimensional construct in women with suspected endometriosis. Thus, the calculation of the total sum scores might not be indicated or proven on a case-by-case basis. No association was found between the expected improvement and the side effects of laparoscopy in all patients. Four distinct clusters described as 'positive', 'no pain, no gain', 'diminished', and 'uniform' were identified. In line with our second hypothesis, the treatment expectation measurements TEX-Q and GEEE correlated higher with each other in terms of convergent validity compared to other psychological measurements assessed in terms of discriminant validity. This applies to the equivalent dimensions of both. Additionally, the expected improvement and worsening of disability (GEEE) and the severity of the five most prevalent endometriosis-related symptoms (NRS) led to medium-to-high positive associations of equivalent dimensions. Both scales used similar wording but focused on different constructs. Contrary to our second hypothesis of convergent validity, correlations of expected disability after laparoscopy measured by the expected PDI and most other measurements of treatment expectations were non-significant to small. However, none of the investigated measurements of treatment expectation correlated to such an extent that they measured exactly the same construct.

## Comparison with existing literature

Treatment expectations measured with the TEX-Q of women with suspected endometriosis undergoing laparoscopy were slightly lower than those of the validation sample<sup>28</sup> for the total score and all dimensions, despite equal scores for the expected 'negative impact' of treatment. This contradicts the findings of more positive treatment expectations in surgical treatment samples undergoing endocrine or bariatric surgery than in psychosomatic samples.<sup>28</sup> Intercorrelations of dimensions were similar in both studies despite of higher correlations between the positive dimensions 'treatment benefit' and 'positive impact' (r=.75 in our sample vs. r=.48 in the validation sample<sup>28</sup>) and 'positive impact' and 'process' (r=.42 vs. r=.15). Moreover, the dimension 'behavioral control' was significantly correlated to 'treatment benefit' (r=.24), 'negative impact' (r=.12), and 'process' (r=.31) in women with suspected endometriosis but not in the validation sample. Correlations of the TEX-Q and other psychological constructs were small to non-significant in our study and the validation study, <sup>28</sup> indicating discriminant validity. Interestingly, participants in our study who reported more depressive symptoms and anxiety expected more adverse events. More unfavorable expectations, that is, lower expected treatment benefit, higher negative impact, and a worse expected process, were also reported in patients with higher depressive and anxiety symptoms in the TEX-Q validation sample.<sup>28</sup> For the other assessed treatment expectation measurements, no validation data are currently available.

In line with Shedden-Mora et al. (2023)<sup>28</sup> and Younger et al. (2012),<sup>24</sup> our findings support the multidimensionality of the construct of treatment expectations: the positive and negative dimensions of treatment expectations were not correlated in these three studies. Overall, the expected side effects of laparoscopy and its improvement were not associated with the results of patients undergoing psychotherapy. <sup>40</sup> This contradicts the 'no pain, no gain' assumption.

So far, research on psychometric properties of measuring treatment expectations <sup>24,28</sup> has used heterogeneous samples including patients undergoing different surgeries, pain, psychosomatic in- or outpatient, or cancer treatment. Differences between patient groups and within patient groups could hardly be detected, but seem promising. Our cluster analysis revealed four distinct clusters in patients with suspected endometriosis that differed in terms of their treatment expectation pattern. These clusters can be described as follows. The first cluster showed a strong positive expectation, followed by no worsening or side effect expectations. For the second cluster, the 'no pain, no gain' assumption seems to be appropriate; these patients expected high side effects of laparoscopy and high improvement. In the third cluster, we found an overall low level of expectations for any category. The fourth cluster had a rather uniform distribution of expectations, with all expectations being of similar strength. Notably, only patients in this cluster patients also expected worsening. The largest cluster was the first with an improvement in expectation dominance, supporting the rather high expectations found in this sample.

## Strengths and limitations

Our findings are based on data from a large sample of women with suspected endometriosis facing laparoscopy in a specialized center for surgical endoscopy and endometriosis, indicating high ecological validity. We assessed four measurements for treatment expectation (TEX-Q, the three items assessing expectations of the GEEE, expected NRS, and expected Pain Disability Index) simultaneously. Consequently, we were able to compare different dimensions, comprehensive questionnaires to single items, and treatment expectation measurements (TEX-Q and GEEE) to established symptom and disability scales (NRS assessing pain and PDI) adapted to the context of expectations. To our knowledge, this is the first study to investigate the treatment expectations in women with suspected endometriosis. Endometriosis is a common chronic disease that is largely underexplored and burdensome, and satisfactory treatment options are not available for many patients.

However, the timeframe of the assessed treatment expectations was not well defined in this study. Women with suspected endometriosis might have different expectations for the short-, medium-, and long-term outcomes after laparoscopy.

We were unable to quantify these potential differences. The order of the measurement instruments was the same for all participants, and sequence effects were possible. To enhance comprehensibility, the survey started with items referring to the past, then to the present, and finally to the future. Consequently, expected disability (expected PDI) and severity of symptoms (expected NRS) were not assessed directly after the preoperative disability and severity of symptoms. Some patients may have forgotten their reference values, leading to imprecise ratings. Finally, some anchors (i.e., the greatest improvement/worsening imaginable) might be understood unequally by participants with suspected endometriosis. Qualitative data from the embedded study module of this cohort study<sup>27</sup> may provide further clarifications.

#### Implications for practice and research

Women with suspected endometriosis are highly burdened and report rather positive expectations concerning laparoscopy, even though laparoscopy is associated with only short-term symptom improvement, <sup>12</sup> and 20–30% do not respond satisfactorily to this treatment. <sup>13,14</sup> Future studies should investigate whether more positive expectations lead to better treatment outcomes or disappointment due to overly optimistic expectations. Based on knowledge of other medical conditions <sup>1</sup> and surgical treatments, <sup>4,5</sup> positive yet realistic expectations (placebo effect) and reduced negative expectations (nocebo effect) may improve treatment outcomes after laparoscopy.

Our results emphasize that treatment expectations should be measured both multidimensionally and comprehensively. The expected improvement and worsening are not the anchors of a single scale. The dimension and linguistic formulation seem more important than the exactly chosen construct, that is, the disability or severity of symptoms. None of the included measurements comprised different time frames. In future research, it is important to define time frames for treatment expectation measurements, especially for patients undergoing surgery, such as laparoscopy. To generalize our findings across patient groups, provide specific recommendations for patients with symptoms of endometriosis, and classify expectations in terms of being real, that is, plausible or ideal <sup>17,19</sup> future qualitative and quantitative longitudinal research is needed. The four clusters of expectations need to be replicated and confirmed because of their high relevance to clinical implications. Individualized interventions could target at expectation manipulation such as focusing on i) side effect management in the 'no pain, no gain' and 'uniform' expectation clusters, ii) increased positive expectations fostering the placebo effect in the 'diminished' cluster, iii) decreased negative expectations reducing the nocebo effect and/or appropriate coping mechanisms, and iv) positive but yet realistic expectations in the 'positive' cluster.

As we found differences between the measurements of treatment expectations and dimensions, it is very interesting whether clinical outcomes may also differ depending on the chosen measurement. In the past, patients' expectations were often assessed heterogeneously without conceptual standardization and psychometric evaluation. <sup>17</sup> Consequently, the selection of measurements should be carefully considered and either tailored to the respective theory-based primary outcome or defined in a general way to be applicable across treatments.

## Ethical approval

Ethical approval was obtained from Psychotherapeutenkammer Hamburg, Germany (ROXWELL-2021-HH, 25<sup>th</sup> of June 2021). This trial was registered at ClinicalTrials.gov (ID: NCT05019612).

#### Data availability

PsychArchives: Dataset for: Assessment of treatment expectations in women with suspected endometriosis: A psychometric analysis, https://doi.org/10.23668/psycharchives.13585.<sup>39</sup>

Data are available under the terms of the Creative Commons Attribution 4.0 International license (CC-BY-SA 4.0).

#### **Acknowledgements**

We thank all patients who participated in this study for their time and involvement. We thank the staff of the *Frauenklinik* an der Elbe and all the psychology students who worked on the ROXWELL study for their support in the recruitment process.

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