UNIVERSITY of York

This is a repository copy of Comparison of the Factor Structure of the Patient Health Questionnaire for somatic symptoms (PHQ-15) in Germany, the Netherlands, and China:A Transcultural Structural Equation Modeling (SEM) Study.

White Rose Research Online URL for this paper: <u>https://eprints.whiterose.ac.uk/133504/</u>

Version: Published Version

### Article:

Leonhart, R., de Vroege, Lars, Schaefert, Rainer et al. (7 more authors) (2018) Comparison of the Factor Structure of the Patient Health Questionnaire for somatic symptoms (PHQ-15) in Germany, the Netherlands, and China:A Transcultural Structural Equation Modeling (SEM) Study. Frontiers in Psychology. 240. ISSN 1664-1078

https://doi.org/10.3389/fpsyt.2018.00240

#### Reuse

This article is distributed under the terms of the Creative Commons Attribution (CC BY) licence. This licence allows you to distribute, remix, tweak, and build upon the work, even commercially, as long as you credit the authors for the original work. More information and the full terms of the licence here: https://creativecommons.org/licenses/

#### Takedown

If you consider content in White Rose Research Online to be in breach of UK law, please notify us by emailing eprints@whiterose.ac.uk including the URL of the record and the reason for the withdrawal request.



eprints@whiterose.ac.uk https://eprints.whiterose.ac.uk/





## Comparison of the Factor Structure of the Patient Health Questionnaire for Somatic Symptoms (PHQ-15) in Germany, the Netherlands, and China. A Transcultural Structural Equation Modeling (SEM) Study

Rainer Leonhart<sup>1†</sup>, Lars de Vroege<sup>2,3\*†</sup>, Lan Zhang<sup>4</sup>, Yang Liu<sup>4</sup>, Zaiquan Dong<sup>4</sup>, Rainer Schaefert<sup>5,6,7</sup>, Sandra Nolte<sup>8,9</sup>, Felix Fischer<sup>8</sup>, Kurt Fritzsche<sup>10</sup> and Christina M. van der Feltz-Cornelis<sup>2,3</sup>

OPEN ACCESS

#### Edited by:

Martina De Zwaan, Hannover Medical School, Germany

#### Reviewed by:

Urs Nater, Universität Wien, Austria Jan Ilhan Kizilhan, University of Duhok, Iraq

#### \*Correspondence:

Lars de Vroege I.devroege@ggzbreburg.nl

<sup>†</sup>These authors have contributed equally to this work.

#### Specialty section:

This article was submitted to Psychosomatic Medicine, a section of the journal Frontiers in Psychiatry

Received: 28 March 2018 Accepted: 17 May 2018 Published: 26 June 2018

#### Citation:

Leonhart R, de Vroege L, Zhang L, Liu Y, Dong Z, Schaefert R, Nolte S, Fischer F, Fritzsche K and van der Feltz-Cornelis CM (2018) Comparison of the Factor Structure of the Patient Health Questionnaire for Somatic Symptoms (PHQ-15) in Germany, the Netherlands, and China. A Transcultural Structural Equation Modeling (SEM) Study. Front. Psychiatry 9:240. doi: 10.3389/fpsyt.2018.00240 <sup>1</sup> Department Social Psychology and Methodology, Institute of Psychology, University of Freiburg, Freiburg, Germany,
<sup>2</sup> Clinical Centre of Excellence for Body Mind and Health, GGz Breburg, Tilburg, Netherlands, <sup>3</sup> Department Tranzo, Tilburg School of Social and Behavioral Sciences, Tilburg University, Tilburg, Netherlands, <sup>4</sup> Mental Health Center, West China Hospital of Sichuan University, Chengdu, China, <sup>5</sup> Division of Internal Medicine, Department of Psychosomatic Medicine, University Hospital Basel, Basel, Switzerland, <sup>6</sup> Department of General Internal Medicine and Psychosomatics, University of Heidelberg, Germany, <sup>7</sup> Faculty of Medicine, University of Basel, Basel, Switzerland, <sup>8</sup> Department of Psychosomatic Medicine, Conter for Internal Medicine and Dermatology, Charité - Universitätsmedizin Berlin, corporate member of Freie Universität Berlin, Humboldt-Universität zu Berlin, and Berlin Institute of Health, Berlin, Germany, <sup>9</sup> Population Health Strategic Research Centre, School of Health and Social Development, Deakin University, Burwood, VIC, Australia, <sup>10</sup> Department of Psychosomatic Medicine and Psychotherapy, Faculty of Medicine, Medical Center - University of Freiburg, Germany

**Background:** Persistent somatic symptoms are associated with psychological distress, impaired function, and medical help-seeking behavior. The Patient Health Questionnaire (PHQ)-15 is used as a screening instrument for somatization and as a monitoring instrument for somatic symptom severity. A bifactorial model has been described, with one general factor and four orthogonal specific symptom factors. The objective of the present study was to assess and to clarify the factor structure of the PHQ-15 within and between different countries in Western Europe and China.

**Method:** Cross-sectional secondary data analysis performed in three patient data samples from two Western European countries (Germany N = 2,517, the Netherlands N = 456) and from China (N = 1,329). Confirmatory factor analyses (CFA), and structural equation modeling (SEM) analysis were performed.

**Results:** The general factor is found in every sample. However, although the outcomes of the PHQ-15 estimate severity of somatic symptoms in different facets, these subscales may have different meanings in the European and Chinese setting. Replication of the factorial structure was possible in the German and Dutch datasets but not in the dataset from China. For the Chinese dataset, a bifactorial model with a different structure for the cardiopulmonary factor is suggested. The PHQ-15 could discern somatization from anxiety and depression within the three samples.

**Conclusion:** The PHQ-15 is a valid questionnaire that can discern somatization from anxiety and depression within different cultures like Europe or China. It can be fitted to

June 2018 | Volume 9 | Article 240

a bifactorial model for categorical data, however, the model can only be recommended for use of the general factor. Application of the orthogonal subscales in non-European samples is not corroborated by the results. The differences cannot be ascribed to differences in health care settings or by differences in concomitant depression or anxiety but instead, a cultural factor involving concepts of disease may play a role in this as they may play a role in the translation of the questionnaire. Further research is needed to explore this, and replication studies are needed regarding the factorial structure of the PHQ-15 in China.

Keywords: somatic symptoms, patient health questionnaire-15, factor structure, structural equation modeling (SEM), transcultural

## BACKGROUND

## Somatic Symptoms, Emotional Distress, and Disability

In everyday life, somatic symptoms are common causes of outpatient medical visits (1, 2). Patients with multiple distressing somatic symptoms present themselves in a variety of health care settings, such as primary, secondary and tertiary patientcenters (3-6). Poor self-rated health of patients was found to be associated with multiple somatic symptoms in Europe and in China (7-10). The number of somatic symptoms correlates well with impaired function and medical help-seeking behavior even after controlling for mental disorders (5). The number of somatic symptoms differs only slightly between patients with somatic symptoms explained by a medical disease and patients with unexplained somatic symptoms (11, 12). This indicates that the suffering of a patient with medically unexplained somatic symptoms should be taken seriously. In Western countries, a high number of somatic symptoms is associated with higher psychological distress, more functional impairment, higher disability, more health care utilization and a reduced quality of life (13-15). Given the large burden of somatic symptoms, especially unexplained symptoms that are currently not taken sufficiently seriously, the assessment of somatic symptoms and the analysis of the different facets of somatic symptoms is a useful and necessary part of every medical diagnostic process.

#### The Patient Health Questionnaire (PHQ)-15

The Patient Health Questionnaire (PHQ)-15 (16) is a short, practical self-rating instrument for the screening of somatic symptoms. The PHQ-15 has also been suggested by the DSM-5 Workgroup on Somatic Symptom Disorders (SSD) as a measurement tool of somatic symptom severity for the classification of SSD (17, 18). The PHQ-15 was initially validated in primary care and a general hospital setting in the USA (16). Although usually the sum score is simply used as a measure for symptom load, several studies (16, 19–21) suggest that the different somatic symptoms in the PHQ-15 can be divided and bundled into four groups: cardiopulmonary, gastrointestinal, pain, and fatigue/general symptoms.

Witthöft et al. (22) assessed the underlying structure of the PHQ-15 in two datasets of college students (Germany, N = 1,520; Switzerland, N = 3,053). They conducted a

confirmatory factor analysis assuming a bifactor model (1 general and 4 orthogonal specific symptom factors; gastrointestinal, fatigue, cardiopulmonary, and pain symptoms). Correlations with general and lower-order latent factors of depression, general somatic symptom distress, and health anxiety were found. These factors explain up to nearly 70% of the variance in the general somatic symptom factor. However, it remains to be shown if this structure also applies to other samples. The original validation of the PHQ-15 was in general hospital patients and primary care patients (16), but not in patients in specialty mental health institutions, so it would be useful to explore the factor structure in other health care settings such as the specialty mental health care setting.

Also, it is unclear whether the factor structure of the PHQ-15 questionnaire is comparable between samples from Europe and samples from other continents and cultures such as Asia, in particular China, since the cultural background shapes the interpretation of somatic symptoms and thereby influences an individual's illness perception and illness behavior (23, 24). In the literature, the influence of culture on experiencing somatic as well as psychosomatic symptoms has been explored. Evidence exists that cultural and personal explanatory models can contribute to the symptomatology of medically unexplained symptoms (25). Efforts have been made to make epidemiological research into this domain possible (26) and the need for appropriate instruments to assess physical symptoms in the context of distress in Asian cultures has been expressed (27).

Historically, there has been a popular belief that Asians manifest a lower prevalence of mood and anxiety disorders than their Western counterparts because they are more prone to experiencing and manifesting distress via somatic pathways (28–30). Among Chinese patients receiving psychiatric services, somatic symptoms such as pain, insomnia, and fatigue have been associated with depressive and anxiety disorders (31).

The PHQ-15 has been translated into many other languages and has been examined in samples from many other countries, e.g., Saudi Arabia, Spain, and Korea (32–34). This offers the potential for comparisons between different ethnic, cultural or geographical groups. However, so far, no studies have explored the factor structure in the context of different health care settings and West European vs. Chinese culture.

Another point of discussion concerns somatization. Somatization is defined as the tendency to experience and

to express physical distress and symptoms that cannot be explained by pathological findings, to attribute them to a medical condition, and to seek medical care for them (35). It has also been described as a term related to our conceptualisation of disease, that is, to the tendency of doctors to attribute symptoms either to physically explainable medical conditions, or as medically unexplained symptoms (36). It has been suggested that physical symptoms occurring in the context of somatization may also concur with depression or anxiety (37). Hence, somatization, as assessed by PHQ-15, might be related to anxiety or depression as assessed by GAD-7 and PHQ-9, as somatization, anxiety, and depression are supposed to be related. It was hypothesized that this might be the case in China, as somatization has been assumed to be more common in China as a manifestation of underlying anxiety and depression (38). Hence, the association between scores on the PHQ-15 and GAD-7 and PHQ-9 should be explored in these different cultures.

## **Rationale and Aims of the Study**

In this study, we compare patients with SSD including all kinds of psychosomatic illnesses in a convenience sample from three different countries, namely Germany, the Netherlands, and China, that enabled us to compare three different health care settings, namely the general hospital inpatient psychosomatic setting (Germany), the specialty mental health outpatient setting providing treatment for SSRD patients with high complexity levels in terms of diagnostic problems, treatment issues and social challenges (8), in the Netherlands; and the general hospital (China) for SSD. Finally, this way we could compare two different cultures, namely the Western European and the Chinese culture.

The objective of the present study was to assess and to clarify the factor structure of the PHQ-15 within and between different countries.

We aimed to answer the following research questions:

- (1) Can the proposed bifactor model of Witthöft et al. (22) be replicated in three different study populations?
- (2) If not—are there more appropriate models within the three study populations?
- (3) Are there indications for cultural differences?
- (4) Is there an association between PHQ-9 and GAD-7 scores and the Witthöft factorial structure of the PHQ-15? More specifically, is there an association between PHQ-9 and GAD-7 scores and the potential general factor and/or the specific factors of the PHQ-15?

## **METHODS**

## **Study Design**

This is a cross-sectional, secondary analysis of data collected within several research projects or routine clinical care in the three countries. In Germany, the data were collected as part of routine clinical care. In China, the data were collected as part of a cross sectional study (39) in the general hospital setting. In the Netherlands, the data were collected as part of routine clinical care in the Clinical Centre of Excellence of Body Mind and Health (8, 40–42).

## Assessment Instruments

#### PHQ-15

The PHQ-15 is a self-administered somatic symptoms subscale derived from the full PHQ (14, 43). The PHQ-15 was originally validated by Spitzer et al. (16). It includes 15 prevalent somatic symptoms that represent over 90% of the symptoms observed in primary care (16). The patients are asked to rate the severity of their symptoms during the previous 4 weeks on a 3-point scale as either 0 ("not bothered at all"), 1 ("bothered a little") or 2 ("bothered a lot"). The questionnaire demonstrated good internal consistency [Cronbach's alpha = 0.80; (16)] and high relevance to symptoms. It is available in multiple languages.

#### Germany

The German translation of the PHQ-15 has been shown to have sound psychometric properties (44) and reference data from the general population is available (45). The PHQ-15, as well as the PHQ-9 and the GAD-7, were collected within the routine psychometric assessment of outpatients in the Department of Psychosomatic Medicine. Data analyzed here has been collected between January 2005 and February 2009. If patients were assessed multiple times, we included the first occasion only. Clinical diagnoses were made by trained physicians and psychologists. Along with questionnaires and clinical diagnoses, we collected sociodemographic data.

#### China

The validity and reliability of the Chinese version of the PHQ-15 (46) were examined in the general population of Hong Kong. The Hong Kong version of the PHQ-15 exhibited satisfactory internal consistency (Cronbach's alpha = 0.79) and stable 1-month test-retest reliability. Somatic symptom severity was positively associated with functional impairment and health service use. In Mainland China, the validity and reliability of the PHQ-15 were tested in the outpatient clinics of general hospitals in Shanghai (47). Cronbach's alpha was 0.73, and the test-retest reliability coefficient was 0.75 (10). There were moderate positive correlations between the PHQ-15 score and anxiety and depression values.

### Translation of the PHQ-15

Three native Chinese speakers who resided in Germany updated the Chinese version of the PHQ-15 taking findings from recent research into account. The translation was based on the former Chinese version, which was translated from English to Mandarin (47). The differences to this former version include changes in item 1 and item 8: We changed "stomach pain" (胃痛或肚痛) in item 1 to "stomach and abdominal pain" (胃痛, in accordance to the suggestions of Lee, Ma (46). In item 8, "fainting spells" was replaced with "brief fainting" [短倒 (Cantonese), 短时间晕倒 (Mandarin)]. The Mandarin version of the PHQ-15 used in this study is available in the Appendix (Supplementary Material).

The data analyzed here were taken from a cross-sectional survey under routine clinical conditions. Participants in this study were inpatients recruited from different departments (e.g., oncology, cardiology, respiratory medicine, rehabilitation, geriatrics and gerontology, general practice, pain management, thyroid and breast surgery, rheumatology, and hepatic surgery). The validation of the PHQ-15 is a component of a larger project investigating the prevalence and recognition of inpatients with emotional distress and their treatment needs at a general hospital.

## **Ethics Statement**

All patients were informed at intake that the Patient Related Outcome Monitoring (PROM) data pertaining to their treatment could be used on an anonymous basis for research, and were provided the opportunity to object to that during intake or anytime during treatment. PROM data of patients objecting to this were not used in this study. The study was approved by the ethics committees of the Shanghai Dong Fang hospital (XZ), the University Medical Centre Freiburg (KF), and GGz Breburg (CFC).

## The Netherlands

In the Netherlands, the PHQ-15 was applied in several research projects in primary care (48, 49), occupational health care (50), and the general hospital setting (51). In primary care, the PHQ-15 showed limited results in terms of validity (52) as it was found to have a tendency to be falsely positive for somatoform disorder, as it does not discern unexplained physical symptoms from explained physical symptoms. In occupational health care, validity was also limited both in terms of sensitivity as well as in terms of specificity (53). It is currently regularly applied in the intake procedure of the Clinical Centre of Excellence for Body, Mind, and Health (Dutch abbreviation: CLGG), which is a specialty mental health institution for patients with somatic symptom and related disorders. Data from this patient group is included in this comparative study. Before intake, patients at CLGG receive questionnaires using Routine Outcome Monitoring. Data for this study were collected during 2010 and 2016. A total of 465 patients filled in the PHQ-15, GAD-7, and PHQ-9.

## Depression Scale PHQ-9

This instrument assesses each of the nine DSM-IV depression criteria on a scale of "0" (not at all) to "3" (nearly every day) (13). The PHQ-9 demonstrated acceptable psychometric properties for the screening of patients with late-life depression. In the Netherlands, the PHQ-9 was validated in the Depression Initiative, a national study aimed at improving mental health care for people with depressive symptoms. It proved to be a feasible and valid instrument to screen for depressive disorder, to assess severity and to assess change in scores over time in the primary care, occupational health care and general hospital outpatient setting. It has also been proven to be feasible and valid for use in electronic monitoring systems in the context of blended mental health care in the occupational health setting. The PHQ-9 is a widely used and valid instrument in the Netherlands as well as in Germany (54–56).

In Chinese, within a primary care setting, this questionnaire showed a sensitivity of 0.86 and a specificity of 0.77 for depression screening (57).

## Anxiety Scale (GAD-7)

A seven-item anxiety scale (GAD-7) was used to assess the severity of generalized anxiety (58). In a German validation study

(59), it has been shown to be a reliable and valid measure of anxiety in the general population. In a study in the Dutch primary Care setting, it showed limited validity as a screener and better validity as a case-finding instrument for general practitioners (60) and it proved to be feasible in the application of collaborative care in the primary care setting. In a Chinese general hospital population, this instrument showed good reliability and good criterion, construct, factorial, and procedural validity (23).

## Statistical Analyses

Using IBM SPSS (24.0) (61), and MPlus (62) 7.4 software, all three samples were analyzed using confirmatory factor analyses (CFA) and structural equation modeling (SEM). As the items of all questionnaires were categorical ordered the analyses were conducted with the robust weighted least squares estimator with a mean and variance adjusted test statistic (WLSMV). Because the chi-square test is sensitive to potentially irrelevant deviations from the implied model structure given large samples, we used established fit measures to evaluate the models. For the absolute fit, we chose the root mean square error of approximation (RMSEA). The comparative fit index (CFI) and the TLI (Tucker-Lewis Index) are reported as incremental fit indices. Following the advice of Hu and Bentler (63) RMSEA values lower than 0.06 and CFI/TLI values higher than 0.95 are considered as indicators of a good model fit. All coefficients (i.e., factor loadings and path weights) are reported as standardized coefficients.

Several CFA models were tested. In a first step, the bifactor model from Witthöft, Fischer (22) was estimated within a multigroup design. Because model fit was only acceptable, alternative models were sought in a second step. In this second step, the bifactor model [see **Figure 1** (Germany), **Figure 2** (Netherlands), and **Figure 3** (China)] was estimated within each group and modification indices were assessed to improve fit indices over abovementioned cutoffs. In a third step, SEMs consisting of measurement models and a structural model was computed in order to determine the size of associations between the latent factors implied by the bifactor model and other relevant constructs (GAD-7 and PHQ-9).

## RESULTS

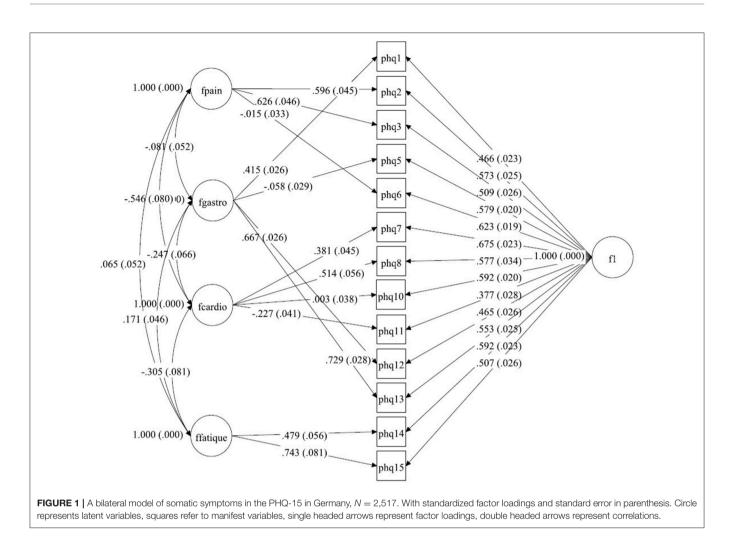
## **Description of the Sample**

Age and gender in the three samples are shown in **Table 1**.

The age and gender are comparable in the dataset from Germany and the Netherlands but the participants from China are significantly older, with a range of up to over 90 years, and have significantly more male participants. This may have to do with the fact that for both Germany and the Netherlands, but not in China, separate health services exist for patients older than 70 years.

The distributions of the PHQ-15 items within the three samples (Germany N = 2,517, China N = 1,329, Netherlands N = 456) are presented in **Table 2.** 

Some items (e.g., PHQ8 [fainting spells] and PHQ11 [pain or problems during sexual intercourse]) have a bottom or floor effects so a metric scale cannot be assumed.



# CFA Models for the Three Samples Taken Together

First, the bifactor model from Witthöft et al. (22) consisting of a general symptom distress factor and four orthogonal symptom-specific factors was estimated within a multigroup design. Our Model showed an acceptable but not excellent model fit in the three samples taken together. The Chi-Square-Test of the model fit was 1850.013 (df = 214, p < 0.0001) with a Chi-Square contribution from the German data of 1081.462, China about 531.294 and the Netherlands 237.257. The RMSEA was 0.073 (90%C.I. = 0.070, 0.076). With a CFI of 0.928 a TLI of 0.921, the first model only has an acceptable fit but not a good model fit.

Hence, a second model was explored assuming that the factors might not be completely independent, allowing for correlations between the four symptom-specific factors to be unequal to zero, in which the four symptom-specific factors would not be orthogonal anymore. This second model was explored over the three groups taken together again.

The Chi-Square-Test of the model fit was 641.186 (df = 46, p < 0.0001). The RMSEA was about 0.055 (90%C.I. = 0.051, 0.059). With a CFI of 0.980, a TLI of 0.965, this model had a better fit but the MPlus results showed that the latent variable covariance

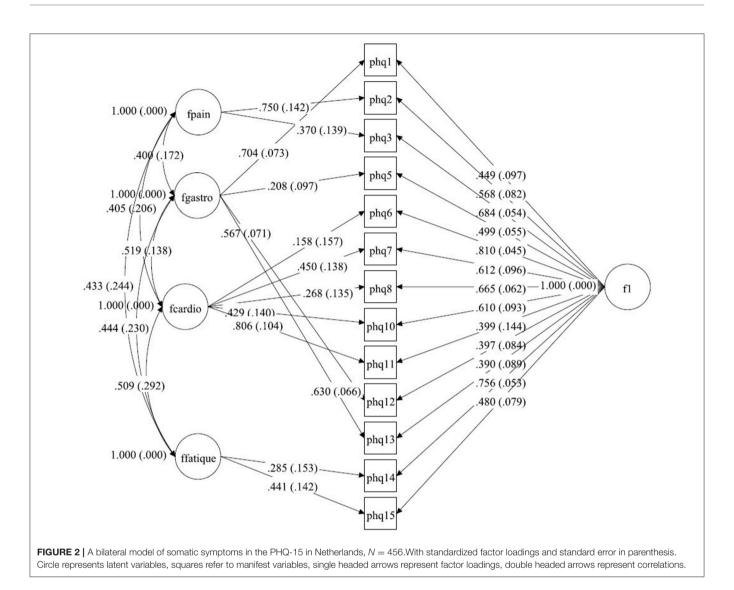
matrix was not positive definite for the Chinese sample, not for the whole model. Hence, a satisfactory fit for the three samples taken together could not be established.

## **CFA Model for the Separate Datasets**

We split the dataset into the three datasets or Germany, China, and the Netherlands, and searched some specific model per country and special sample within these countries.

#### Germany

Starting with the German dataset, we tried a model comparable to Witthöft et al. (22) but allowing for correlations between the four symptom-specific factors. The Chi-Square-Test of the fit for this final German model was 265.638 (df = 46, p < 0.0001). The RMSEA was about 0.044 (90%C.I. = 0.039, 0.049). With a CFI of 0.983, a TLI of 0.971. The model therefore fits in the German dataset. The only difference between Witthöft et al. (22) are the low but essential correlations between the factors. Without these correlations, the model would not fit. The results of this model are shown in **Tables 2a,b**. Almost all correlations between the four factors were significant. Two factors (Pain and Cardio) have a correlation of about -0.546; the others have a lower absolute value. The model is shown in **Figure 1**.



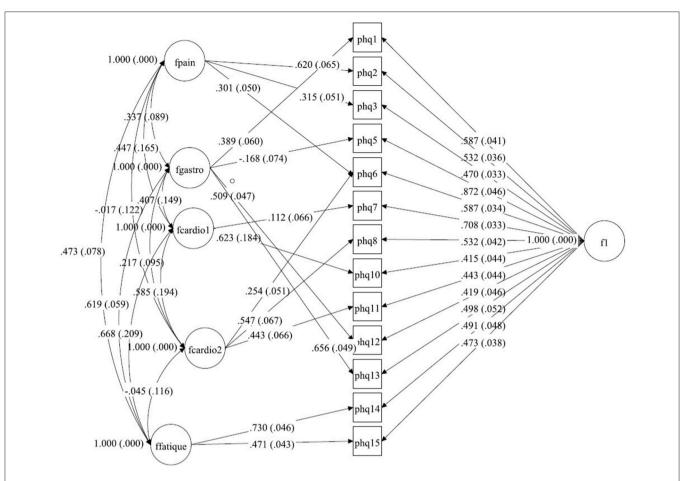
#### The Netherlands

For the dataset from the Netherlands, explorative testing and using the modification indices showed that only a little change for item 6 (chest pain) had to be done (see **Tables 3a,b**). In the original model, this item belongs to painrelated symptoms. For our model, this item fits better to the cardiopulmonary-related symptoms factor. The Chi-Square-Test of the fit for the final Dutch model was 53.624 (df = 46, p = 0.2051). The RMSEA was about 0.019 (90%C.I. = 0.000, 0.038; probability of RMSEA  $\leq 0.05$  is = 0.999). With a CFI of 0.998, a TLI of 0.997. The model is shown in **Figure 2**.

#### China

The structure from Witthöft et al. (22) could not be replicated in the dataset from China. We split the cardiopulmonaryrelated symptoms factor into two factors. These factors were correlated but estimate different latent variables. In the model for this dataset the pain-related symptoms factor includes items PHQ2 (back pain), PHQ3 (pain arm legs), and PHQ6 (chest pain), and the gastrointestinal-related symptoms factor includes items PHQ1 (stomach pain), PHQ5 (headaches), PHQ12 (constipation), and PHQ13 (nausea, gas). The first cardiopulmonary-related symptoms factor only includes items PHQ7 (dizziness) and PHQ10 (short breath), while the second cardiopulmonary-related symptoms factor is measured by PHQ6 (chest pain), PHQ8 (fainting spells), and PHQ11 (pain sexual intercourse) (see **Tables 4a,b**). The fatigue-related symptoms factor includes items PHQ14 and PHQ15. The Chi-Square-Test of the fit for the final Chinese model was 187.779 (df = 41, p < 0.0001). The RMSEA was about 0.052 (90%C.I. = 0.045, 0.060; probability of RMSEA  $\leq 0.05$  is = 0.325). A CFI of 0.977, a TLI of 0.957, show a good fit of the model, as shown in **Figure 3**.

A part of the results from the different models—the thresholds between the points on the scale of the PHQ-15 items—can be interpreted and compared between the different countries. The results in **Table 5** reflect the descriptive in **Table 1** where the



**FIGURE 3** | A bilateral model of somatic symptoms in the PHQ-15 in China, N = 1,329. With standardized factor loadings and standard error in parenthesis. Circle represents latent variables, squares refer to manifest variables, single headed arrows represent factor loadings, double headed arrows represent correlations.

|        |            | Germany | China  | Netherlands | F/Chi2  | df                  | p       | R <sup>2</sup> |
|--------|------------|---------|--------|-------------|---------|---------------------|---------|----------------|
| Total  | Ν          | 2,517   | 1,322  | 456         |         |                     |         |                |
| Age    | Mean       | 42.56   | 53.76  | 41.02       | 259.68  | df1 = 2 df2 = 4,292 | < 0.001 | 0.108          |
|        | SD         | 15.13   | 16.21  | 12.65       |         |                     |         |                |
| Gender | Male (N)   | 782     | 771    | 167         |         |                     |         |                |
|        | (%)        | 31.10%  | 58.10% | 36.60%      | 266.27% | 2                   | < 0.001 |                |
|        | Female (N) | 1,735   | 557    | 289         |         |                     |         |                |
|        | (%)        | 68.90%  | 41.90% | 63.40%      |         |                     |         |                |

TABLE 1 | Demographic characteristics.

participants from the different samples respond differently. This diversity leads to different thresholds in the three samples. They can be deduced from **Table 1** and are shown in the Table below. For example, in the PHQ-15 item 3 the thresholds for both European samples are nearly the same while the thresholds for the sample from China are substantially higher. However, for all used PHQ-15 items there are thresholds, which show that all three points of the used 3-point scale are useful. This is shown in **Table 5**.

### PHQ-9 and GAD-7

We then tested the predictive value of the PHQ-9 and the GAD-7 questionnaires within these models. Within the German and the Chinese sample the PHQ-9 had good prediction values for most PHQ-15 factors while the GAD-7 predicted the general factor and the fatigue factor well. In the sample from the Netherlands, only the general factor is well-predicted by the PHQ-9 and the GAD-7. This is shown in **Table 6**.

#### TABLE 2 | Descriptive statistics for the items of the PHQ15 for all three samples.

|  |                     | Germany<br>(N = 2,517) | China<br>(N = 1,329) | Netherlands<br>(N = 456)   |
|--|---------------------|------------------------|----------------------|--|
| PHQ1                                       | Not bothered at all | 47.9 % 1,206           | 69.0% 917            | 39.1% 175  |
| stomach pain                               | Bothered a little   | 25.9% 651              | 25.3% 336            | 42.6% 191  |
|  | Bothered a lot      | 26.2% 660              | 5.7% 76              | 18.3% 82   |
| PHQ2                                       | Not bothered at all | 30.1% 757              | 55.2% 733            | 29.6% 135  |
| back pain                                  | Bothered a little   | 29.6% 746              | 36.7% 488            | 40.8% 186  |
|  | Bothered a lot      | 40.3% 1,014            | 8.1% 108             | 29.6% 135  |
| PHQ3                                       | Not bothered at all | 32.2% 811              | 52.9% 703            | 26.5% 121  |
| oain in your arms, legs, or joint          | Bothered a little   | 28.3% 712              | 34.5% 458            | 36.8% 168  |
|  | Bothered a lot      | 39.5% 994              | 12.6% 168            | 36.6% 167  |
| PHQ5                                       | Not bothered at all | 32.3% 814              | 65.4% 869            | 66.9% 305  |
| headaches                                  | Bothered a little   | 37.1% 935              | 30.1% 400            | 24.1% 110  |
|  | Bothered a lot      | 30.5% 768              | 4.5% 60              | 9.0% 41  |
| PHQ6                                       | Not bothered at all | 53.4% 1,344            | 64.0% 850            | 32.5% 148  |
| chest pain                                 | Bothered a little   | 27.7% 698              | 30.8% 409            | 43.2% 197  |
|  | Bothered a lot      | 18.9% 650              | 5.3% 70              | 24.3% 111  |
| PHQ7                                       | Not bothered at all | 36.2% 912              | 57.0% 758            | 57.0% 260  |
| dizziness                                  | Bothered a little   | 37.9% 955              | 38.3% 509            | 30.7% 140  |
|  | Bothered a lot      | 25.8% 650              | 4.7% 62              | 12.3% 56   |
| PHQ8                                       | Not bothered at all | 84.3% 2,121            | 83.4% 1,109          | 41.0% 187  |
| ainting spells                             | Bothered a little   | 10.2% 257              | 13.2% 175            | 40.4% 184  |
|  | Bothered a lot      | 5.5% 139               | 3.4% 45              | 18.6% 85   |
| PHQ10                                      | Not bothered at all | 46.0% 1,158            | 67.0% 891            | 50.4% 230  |
| shortness of breath                        | Bothered a little   | 31.9% 804              | 25.7% 342            | 36.2% 165  |
|  | Bothered a lot      | 22.1% 555              | 7.2% 96              | 13.4% 61   |
| PHQ11                                      | Not bothered at all | 69.9% 1,751            | 82.7% 1,099          | 38.2% 174  |
| pain or problems during sexual intercourse | Bothered a little   | 13.7% 345              | 13.8% 184            | 45.0% 205  |
|  | Bothered a lot      | 16.7% 421              | 3.5% 46              | 16.9% 77   |
| PHQ12                                      | Not bothered at all | 41.0% 1,033            | 50.6% 672            | 37.7% 172  |
| constipation, loose bowels, or diarrhea    | Bothered a little   | 31.1% 783              | 39.3% 522            | 43.4% 198  |
|  | Bothered a lot      | 27.9% 701              | 10.2% 135            | 18.9% 86   |
| PHQ13                                      | Not bothered at all | 34.1% 858              | 51.5% 685            | 20.8% 95   |
| nausea, gas, or indigestion                | Bothered a little   | 32.7% 822              | 39.3% 522            | 59.2% 270  |
|  | Bothered a lot      | 33.3% 837              | 9.2% 122             | 42.6% 191<br>18.3% 82<br>29.6% 135<br>40.8% 186<br>29.6% 135<br>26.5% 121<br>36.8% 168<br>36.6% 167<br>66.9% 305<br>24.1% 110<br>9.0% 41<br>32.5% 148<br>43.2% 197<br>24.3% 111<br>57.0% 260<br>30.7% 140<br>12.3% 56<br>41.0% 187<br>40.4% 184<br>18.6% 85<br>50.4% 205<br>16.9% 77<br>37.7% 172<br>43.4% 61<br>38.2% 174<br>45.0% 205<br>16.9% 77<br>37.7% 172<br>43.4% 198<br>18.9% 86<br>20.8% 95<br>59.2% 270<br>20.0% 91<br>22.4% 102<br>41.7% 190<br>36.0% 164<br>7.0% 19 |
| PHQ14                                      | Not bothered at all | 8.6% 216               | 36.0% 479            | 22.4% 102  |
| eeling tired or having low energy          | Bothered a little   | 28.6% 720              | 50.8% 675            | 41.7% 190  |
|  | Bothered a lot      | 62.8% 1,581            | 13.2% 175            | 36.0% 164  |
| PHQ15                                      | Not bothered at all | 12.5% 314              | 43.5% 578            | 7.0% 19  |
| trouble sleeping                           | Bothered a little   | 25.9% 651              | 40.8% 542            | 18.1% 49   |
|  | Bothered a lot      | 61.7% 1,552            | 15.7% 209            | 74.8% 202  |

Only three categories of possible answers exist within the PHQ15.

## DISCUSSION

This study examines the factor structure of the PHQ-15 in a cross country and cross cultural setting involving China, with a total of 4302 participants. It shows that the factor structure found by Witthöft et al. (22) consisting of a general symptom distress factor and four orthogonal symptom-specific factors (pain-, gastrointestinal-, cardiopulmonary-, and fatigue-related symptoms) does not fit the full sample consisting of all three study populations.

An analysis per sample showed similarities with the Witthöft model in the German and the Dutch population. In the German sample the model of Witthöft could be replicated if low but essential correlations between the factors were allowed. We find a comparable but different model for the dataset from the Netherlands. The Dutch sample showed an excellent fit with the original model if item 6 was adapted belonging to pain-related symptoms toward fitting this item to the cardiopulmonaryrelated symptoms factor.

However, the Witthöft model could not be replicated in the Chinese sample. There were differences in pain, in the sense that the cardiovascular item chest pain was mostly reported in the context of general pain symptoms. Also, items like fainting, chest pain and problems with sexual intercourse were related, contrary to the European samples. Hence the Chinese sample seems to have different properties compared to the European samples. This may be due to epidemiological differences, such as that cardiovascular symptoms. But it might also be associated with cultural differences in body and illness perception, or in scripting, as described by Kleinman (64); a script concerning a specific clustering of symptoms may be shared by patients and Chinese

#### Table 2a | Fitting solution in the German sample.

|       | General Factor (S.E.) | Factor PAIN (S.E.) | Factor GASTRO (S.E.) | Factor CARDIO (S.E.) | Factor FATIGUE (S.E. |
|-------|-----------------------|--------------------|----------------------|----------------------|----------------------|
| PHQ1  | 0.466 (0.023)         |                    | 0.415 (0.026)        |                      |                      |
| PHQ2  | 0.573 (0.025)         | 0.596 (0.045)      |                      |                      |                      |
| PHQ3  | 0.509 (0.026)         | 0.626 (0.046)      |                      |                      |                      |
| PHQ5  | 0.579 (0.020)         |                    | -0.058 (0.029)       |                      |                      |
| PHQ6  | 0.623 (0.019)         | -0.015 (0.033)     |                      |                      |                      |
| PHQ7  | 0.675 (0.023)         |                    |                      | 0.381 (0.045)        |                      |
| PHQ8  | 0.577 (0.034)         |                    |                      | 0.514 (0.056)        |                      |
| PHQ10 | 0.592 (0.020)         |                    |                      | 0.003 (0.038)        |                      |
| PHQ11 | 0.377 (0.028)         |                    |                      | -0.227 (0.041)       |                      |
| PHQ12 | 0.465 (0.026)         |                    | 0.667 (0.026)        |                      |                      |
| PHQ13 | 0.553 (0.025)         |                    | 0.729 (0.028)        |                      |                      |
| PHQ14 | 0.592 (0.023)         |                    |                      |                      | 0.479 (0.056)        |
| PHQ15 | 0.507 (0.026)         |                    |                      |                      | 0.743 (0.081)        |

Table 2b | Correlations between the factors within the German sample.

|               | Factor GASTRO            | Factor CARDIO            | Factor FATIGUE                 |
|---------------|--------------------------|--------------------------|--------------------------------|
| Factor PAIN   | -0.081 (0.052) p = 0.118 | -0.546 (0.080) p < 0.001 | 0.065 (0.052) p = 0.209        |
| Factor GASTRO |                          | -0.247 (0.066) p < 0.001 | 0.171 (0.046) <i>p</i> < 0.001 |
| Factor CARDIO |                          |                          | -0.305 (0.081) p < 0.001       |

Table 3a | Fitting solution in the sample from the Netherlands.

|       | General Factor (S.E.) | Factor PAIN (S.E.) | Factor GASTRO (S.E.) | Factor CARDIO (S.E.) | Factor FATIGUE (S.E. |
|-------|-----------------------|--------------------|----------------------|----------------------|----------------------|
| PHQ1  | 0.449 (0.097)         |                    | 0.704 (0.073)        |                      |                      |
| PHQ2  | 0.568 (0.082)         | 0.750 (0.142)      |                      |                      |                      |
| PHQ3  | 0.684 (0.054)         | 0.370 (0.139)      |                      |                      |                      |
| PHQ5  | 0.499 (0.055)         |                    | 0.208 (0.097)        |                      |                      |
| PHQ6  | 0.810 (0.045)         |                    |                      | 0.158 (0.157)        |                      |
| PHQ7  | 0.612 (0.096)         |                    |                      | 0.450 (0.138)        |                      |
| PHQ8  | 0.665 (0.062)         |                    |                      | 0.268 (0.135)        |                      |
| PHQ10 | 0.610 (0.093)         |                    |                      | 0.429 (0.140)        |                      |
| PHQ11 | 0.339 (0.144)         |                    |                      |                      |                      |
| PHQ12 | 0.397 (0.084)         |                    | 0.567 (0.071)        |                      |                      |
| PHQ13 | 0.390 (0.089)         |                    | 0.630 (0.066)        |                      |                      |
| PHQ14 | 0.756 (0.053)         |                    |                      |                      | 0.285 (0.153)        |
| PHQ15 | 0.480 (0.079)         |                    |                      |                      | 0.441 (0.142)        |

Table 3b | Correlations between the factors within the sample from the Netherlands.

|         | Factor GASTRO           | FCARDIO                 | FFATIGUE                     |
|---------|-------------------------|-------------------------|------------------------------|
| FPAIN   | 0.400 (0.172) p = 0.020 | 0.405 (0.206) p = 0.049 | 0.433 (0.244) p = 0.076      |
| FGASTRO |                         | 0.519 (0.138) p < 0.001 | 0.444 (0.230) p = 0.054      |
| FCARDIO |                         |                         | 0.509 (0.292) $\rho = 0.082$ |

mental health professionals and this might be one of such scripts. Ryder and Chentsova-Dutton (65) describe so-called "scripts"as ... "organized units of culturally salient knowledge, such as knowledge about the ways in which one communicates distress... based on observation as well as formal learning. Scripts serve as mechanisms for rapid, automatic retrieval of information and recognition of patterns. ...Once scripts are enacted they are observable by others as behavior and become elements of the

|       | General Factor (S.E.) | Factor PAIN (S.E.) | Factor GASTRO (S.E.) | Factor CARDIO1 (S.E.) | Factor CARDIO2 (S.E.) | Factor FATIGUE (S.E |
|-------|-----------------------|--------------------|----------------------|-----------------------|-----------------------|---------------------|
| PHQ1  | 0.587 (0.041)         |                    | 0.389 (0.060)        |                       |                       |                     |
| PHQ2  | 0.532 (0.036)         | 0.620 (0.065)      |                      |                       |                       |                     |
| PHQ3  | 0.470 (0.033)         | 0.315 (0.051)      |                      |                       |                       |                     |
| PHQ5  | 0.872 (0.046)         |                    | -0.168 (0.074)       |                       |                       |                     |
| PHQ6  | 0.587 (0.034)         | 0.301 (0.050)      |                      |                       | 0.254 (0.051)         |                     |
| PHQ7  | 0.708 (0.033)         |                    |                      | 0.112 (0.066)         |                       |                     |
| PHQ8  | 0.532 (0.042)         |                    |                      |                       | 0.547 (0.067)         |                     |
| PHQ10 | 0.415 (0.044)         |                    |                      | 0.623 (0.184)         |                       |                     |
| PHQ11 | 0.443 (0.044)         |                    |                      |                       | 0.443 (0.066)         |                     |
| PHQ12 | 0.419 (0.046)         |                    | 0.509 (0.047)        |                       |                       |                     |
| PHQ13 | 0.498 (0.052)         |                    | 0.656 (0.049)        |                       |                       |                     |
| PHQ14 | 0.491 (0.048)         |                    |                      |                       |                       | 0.730 (0.046)       |
| PHQ15 | 0.473 (0.038)         |                    |                      |                       |                       | 0.471 (0.043        |

Table 4a | Fitting solution in the Chinese sample.

Table 4b | Correlations between the factors within the Chinese sample.

|                | Factor GASTRO           | Factor CARDIO1                 | Factor CARDIO2           | Factor FATIGUE           |
|----------------|-------------------------|--------------------------------|--------------------------|--------------------------|
| Factor PAIN    | 0.337 (0.089) p < 0.001 | 0.447 (0.165) <i>p</i> = 0.007 | -0.017 (0.122) p = 0.892 | 0.473 (0.078) p < 0.001  |
| Factor GASTRO  |                         | 0.407 (0.149) p = 0.006        | 0.217 (0.095) p = 0.003  | 0.619 (0.059) p < 0.001  |
| Factor CARDIO1 |                         |                                | 0.585(0.194) p = 0.003   | 0.668 (0.209) p = 0.001  |
| Factor CARDIO2 |                         |                                |                          | -0.045 (0.116) p = 0.697 |

**TABLE 5** | Thresholds between the different steps on the 3-point scale of the PHQ-15.

|              | Germany<br>threshold (S.E.) | China threshold<br>(S.E.) | Netherlands<br>threshold (S.E.) |
|--------------|-----------------------------|---------------------------|---------------------------------|
|              |                             |                           | . ,                             |
| PHQ1 1 to 2  | -0.052 (0.025)              | 0.496 (0.036)             | -0.278 (0.060)                  |
| PHQ1 2 to3   | 0.637 (0.027)               | 1.579 (0.056)             | 0.904 (0.069)                   |
| PHQ2 1 to 2  | -0.522 (0.026)              | 0.130 (0.034)             | -0.536 (0.062)                  |
| PHQ2 2 to3   | 0.246 (0.025)               | 1.397 (0.050)             | 0.536 (0.062)                   |
| PHQ3 1 to 2  | -0.462 (0.026)              | 0.073 (0.034)             | -0.627 (0.069)                  |
| PHQ3 2 to3   | 0.267 (0.025)               | 1.144 (0.044)             | 0.342 (0.060)                   |
| PHQ5 1 to 2  | -0.485 (0.026)              | 0.396 (0.035)             | 0.437 (0.061)                   |
| PHQ5 2 to3   | 0.510 (0.026)               | 1.694 (0.060)             | 1.341 (0.083)                   |
| PHQ6 1 to 2  | 0.085 (0.025)               | 0.357 (0.035)             | -0.455 (0.061)                  |
| PHQ6 2 to3   | 0.883 (0.029)               | 1.619 (0.057)             | 0.695 (0.064)                   |
| PHQ7 1 to 2  | -0.352 (0.026)              | 0.177 (0.035)             | 0.177 (0.059)                   |
| PHQ7 2 to3   | 0.649 (0.027)               | 1.678 (0.059)             | 1.161 (0.076)                   |
| PHQ8 1 to 2  | 1.005 (0.030)               | 0.972 (0.041)             | -0.227 (0.059)                  |
| PHQ8 2 to3   | 1.596 (0.041)               | 1.827 (0.066)             | 0.891 (0.068)                   |
| PHQ10 1 to 2 | -0.100 (0.025)              | 0.441 (0.036)             | 0.011 (0.059)                   |
| PHQ10 2 to3  | 0.771 (0.028)               | 1.459 (0.052)             | 1.109 (0.074)                   |
| PHQ11 1 to 2 | 0.512 (0.026)               | 0.942 (0.041)             | -0.301 (0.060)                  |
| PHQ11 2 to3  | 0.965 (0.030)               | 1.817 (0.065)             | 0.959 (0.070)                   |
| PHQ12 1 to 2 | -0.226 (0.025)              | 0.014 (0.034)             | -0.313 (0.060)                  |
| PHQ12 2 to3  | 0.587 (0.027)               | 1.237 (0.047)             | 0.883 (0.068)                   |
| PHQ13 1 to 2 | -0.410 (0.026)              | 0.039 (0.034)             | -0.812 (0.066)                  |
| PHQ13 2 to3  | 0.433 (0.026)               | 1.330 (0.048)             | 0.843 (0.067)                   |
| PHQ14 1 to 2 | -1.367 (0.036)              | -0.357 (0.035)            | -0.760 (0.065)                  |
| PHQ14 2 to3  | -0.327 (0.025)              | 1.118 (0.043)             | 0.359 (0.060)                   |
| PHQ15 1 to 2 | -1.152 (0.032)              | -0.164 (0.035)            | -1.473 (0.115)                  |
| PHQ15 2 to3  | -0.297 (0.025)              | 1.006 (0.042)             | -0.669 (0.083)                  |
|              |                             |                           |                                 |

larger cultural context" (66). In their elaboration, they refer to Kleinman and Kleinman who explain the meaning of scripts as potentially influenced by political events such as the Cultural Revolution. In this way, symptoms of distress i.e., depression could have dangerous connotations if expressed as such (67, 68) and thus lead to different scripts for distress (68).

Thus, cultural scripts (66) shape the ways in which people attend and react to particular experiences marked as important in some way. In some cases, pathological loops can form, where attention to a particular symptom can accentuate its severity and give rise to related symptoms' (65).

Another explanation may be fear of stigma; if sexual intercourse would be experienced as problematic, it might be reported in the context of symptoms suggesting cardiovascular disease, and not as standalone. A connection between shame and stigma has been reported for schizophrenia (69). The role of stigma in recognition of mental disorders in China has been explored (70). The literature indicates that higher rates of shyness and other normal interpersonal concerns in Chinese cultural contexts exist (71). Such shame may also play a role in sexual problems if it occurs in the context of unrecognized mental disorder, and thus in the reporting of physical symptoms in SSD.

Regarding a possible association between factor structure and health care settings, this was considered as a possibility because somatic symptoms may be perceived as providing more effective access to health care resources (38), however, this does not seem to be corroborated by the results. The German and Dutch samples seem to be congruent although from different health care settings, namely a general hospital setting and a specialized mental health care setting for SSD. The Chinese sample also

#### **TABLE 6** | Different factors predicted by PHQ-9 and GAD-7.

| Factor  | Germany       |        |               |        | China         |        |               |        | Netherlands   |       |               |       |
|---------|---------------|--------|---------------|--------|---------------|--------|---------------|--------|---------------|-------|---------------|-------|
|         | PHQ-9         | 9      | GAD           |        | PHQ-9         | 9      | GAD-7         | 7      | PHQ-9         |       | GAD-7         | ,     |
|         | Est. (SE)     | р      | Est. (SE)     | р     | Est. (SE)     | Р     |
| General | 0.348 (0.027) | <0.001 | 0.433 (0.029) | <0.001 | 0.168 (0.043) | <0.001 | 0.211 (0.038) | <0.001 | 0.178 (0.059) | 0.002 | 0.167 (0.061) | 0.006 |
| Pain    | 0.027 (0.020) | 0.171  | 0.019 (0.020) | 0.331  | 0.160 (0.068) | 0.019  | 0.108 (0.061) | 0.080  | 0.000 (0.002) | 0.980 | 0.035 (0.065) | 0.595 |
| Gastro  | 0.057 (0.020) | 0.004  | 0.017 (0.015) | 0.254  | 0.206 (0.046) | <0.001 | 0.093 (0.035) | 0.009  | 0.036 (0.039) | 0.357 | 0.010 (0.023) | 0.648 |
| Cardio  | 0.085 (0.043) | 0.049  | 0.087 (0.057) | 0.123  | 0.076 (0.047) | 0.107  | 0.027 (0.027) | 0.315  | 0.068 (0.056) | 0.224 | 0.055 (0.058) | 0.337 |
| Fatigue | 0.763 (0.030) | <0.001 | 0.886 (0.035) | <0.001 | 0.402 (0.046) | <0.001 | 0.384 (0.047) | <0.001 | 0.036 (0.074) | 0.623 | 0.001 (0.004) | 0.988 |

Annotation: For this comparison we use only the second cardio factor within the sample from China. Significant regression weights are printed in bold.

concerns a general hospital setting, but the Chinese sample shows a different factorial structure from the European samples.

This different factorial structure in the Chinese sample does not seem to be explained by different associations with depression and anxiety either, as both the German and the Chinese sample showed that the PHQ-9 had good predictive values for most PHQ-15 factors while the GAD-7 well-predicted the general factor and the fatigue factor. In contrast, the sample from the Netherlands found the PHQ-9 and the GAD-7 to be only predictive for the PHQ-15 general factor.

The findings of this study suggest that the different factorial structure of the PHQ-15 in China is not to be explained by differences in concomitant depression or anxiety. Cultural factors such as stigma, scripting, or different concepts of disease (36), may play a role in this. Further research is needed to explore this, and replication studies are needed regarding the factorial structure of the PHQ-15 in China. Nevertheless, with this knowledge the PHQ15 can be applied in China, taking this small difference in somatic symptom clusters for China into account. Future studies should also explore to what extend the different factorial structures might bias data analysis when the PHQ-15 is used as a unidimensional measure. Furthermore, they should investigate whether any confounding factors could be identified which drive the differences between cultures.

#### Limitations of the study

It can be considered a limitation of the study that health care settings and culture differences overlap and hence their assessment cannot be separated sufficiently to enable firm conclusions in this matter.

## CONCLUSION

The PHQ-15 is a valid questionnaire that can discern somatization from anxiety and depression within different

## REFERENCES

 Khan AA, Khan A, Harezlak J, Tu W, Kroenke K. Somatic symptoms in primary care: etiology and outcome. *Psychosomatics* (2003) 44:471–8. doi: 10.1176/appi.psy.44.6.471 cultures like Europe or China. The questionnaire can be fitted to a bifactorial model, however, we can only recommend the use of the general factor. Application of the orthogonal subscales in non-European samples is not corroborated by the results. Further research is needed to explore explanations and to replicate the findings of this study.

## AUTHOR CONTRIBUTIONS

Acquisition of the study was by KF. CvdF-C, KF, RL, and LdV designed the study. CvdF-C, LdV, LZ, YL, ZD, RS, SN, FF provided the data. RL, FF, CvdF-C, LdV, and KF designed the analyses and RL performed them. CvdF-C, LdV, and RS contributed to the interpretation of the data. RL, LdV, and CvdF-C wrote the manuscript and LZ, YL, ZD, RS, SN, and FF contributed to the manuscript for the Chinese and German samples. All authors critically revised and approved the final version of the article.

### FUNDING

Data collection in China was supported by the Grant GZ 1155 of the Sino-German Research Centre. Data analyses and writing of the manuscript were supported by Grant GZ 690 awarded by the Centre for Sino-German Research Promotion in Beijing to Kurt Fritzsche. Furthermore, this study was supported by GGz Breburg and Tilburg University, Tilburg, the Netherlands. The article processing charge was funded by the German Research Foundation (DFG) and the University of Freiburg in the funding programme Open Access Publishing.

### SUPPLEMENTARY MATERIAL

The Supplementary Material for this article can be found online at: https://www.frontiersin.org/articles/10.3389/fpsyt. 2018.00240/full#supplementary-material

van Eck van der Sluijs J, ten Have M, Rijnders CA, van Marwijk HW, de Graaf R, van der Feltz-Cornelis CM. Mental health care use in medically unexplained and explained physical symptoms: findings from a general population study. *Neuropsychiatr Dis Treat.* (2016) 12:2063. doi: 10.2147/NDT.S109504

- Kroenke K, Arrington ME, Mangelsdorff AD. The prevalence of symptoms in medical outpatients and the adequacy of therapy. *Arch Intern Med.* (1990) 150:1685–9. doi: 10.1001/archinte.150.8.1685
- Creed F, Tomenson B, Chew-Graham C, Macfarlane G, Davies I, Jackson J, et al. Multiple somatic symptoms predict impaired health status in functional somatic syndromes. *Int J Behav Med.* (2013) 20:194–205. doi: 10.1007/s12529-012-9257-y
- Tomenson B, Essau C, Jacobi F, Ladwig KH, Leiknes KA, Lieb R, et al. Total somatic symptom score as a predictor of health outcome in somatic symptom disorders. *Brit J Psychiatry* (2013) 203:373–80.
- Creed FH, Davies I, Jackson J, Littlewood A, Chew-Graham C, Tomenson B, et al. The epidemiology of multiple somatic symptoms. *J Psychosom Res.* (2012) 72:311–7. doi: 10.1016/j.jpsychores.2012.01.009
- Schaefert R, Höner C, Salm F, Wirsching M, Leonhart R, Yang J, et al. Psychological and behavioral variables associated with the somatic symptom severity of general hospital outpatients in China. *Gen Hosp Psychiatry* (2013) 35:297–303. doi: 10.1016/j.genhosppsych.2012.11.001
- van Eck van der Sluijs J, de Vroege L, van Manen A, Rijnders CT, van der Feltz-Cornelis C. Complexity assessed by the intermed in patients with somatic symptom disorder visiting a specialized outpatient mental health care setting: a cross sectional study complexity of patients with SSD. *Psychosomatics* (2017) 58:427–36. doi: 10.1016/j.psym.2017.02.008
- Zhu C, Ou L, Geng Q, Zhang M, Ye R, Chen J, et al. Association of somatic symptoms with depression and anxiety in clinical patients of general hospitals in Guangzhou, China. *Gen Hosp Psychiatry* (2012) 34:113–20. doi: 10.1016/j.genhosppsych.2011.09.005
- Zhang L, Fritzsche K, Liu Y, Wang J, Huang M, Wang Y, et al. Validation of the Chinese version of the PHQ-15 in a tertiary hospital. *BMC Psychiatry* (2016) 16:89. doi: 10.1186/s12888-016-0798-5
- Kisely S, Simon G. An international study comparing the effect of medically explained and unexplained somatic symptoms on psychosocial outcome. J Psychosom Res. (2006) 60:125–30. doi: 10.1016/j.jpsychores.2005.06.064
- Escobar JL, Cook B, Chen CN, Gara MA, Alegria M, Interian A, et al. Whether medically unexplained or not, three or more concurrent somatic symptoms predict psychopathology and service use in community populations. J Psychosom Res. (2010) 69:1–8. doi: 10.1016/j.jpsychores.2010.01.001
- Kroenke K, Spitzer RL, Williams JB. The PHQ-9: validity of a brief depression severity measure. J Gen Intern Med. (2001) 16:606–13. doi: 10.1046/j.1525-1497.2001.016009606.x
- Kroenke K, Spitzer RL, Williams JB, Löwe B. The patient health questionnaire somatic, anxiety, and depressive symptom scales: a systematic review. *Gen Hosp Psychiatry* (2010) 32:345–59. doi: 10.1016/j.genhosppsych.2010.03.006
- 15. van Eck van der Sluijs J, Ten Have M, Rijnders C, van Marwijk H, de Graaf R, van der Feltz-Cornelis C. Medically unexplained and explained physical symptoms in the general population: association with prevalent and incident mental disorders. *PLoS ONE* (2015) 10:e0123274. doi: 10.1371/journal.pone.0123274
- Kroenke K, Spitzer RL, Williams JB. The PHQ-15: validity of a new measure for evaluating the severity of somatic symptoms. *Psychosom Med.* (2002) 64:258–66. doi: 10.1097/00006842-200203000-00008
- Narrow WE, Clarke DE, Kuramoto SJ, Kraemer HC, Kupfer DJ, Greiner L, et al. DSM-5 field trials in the United States and Canada, Part III: development and reliability testing of a cross-cutting symptom assessment for DSM-5. *Am J Psychiatry* (2013) 170:71–82. doi: 10.1176/appi.ajp.2012.12071000
- American Psychiatric Association. *Diagnostic and Statistical Manual* of *Mental Disorders (DSM-5)*. Washington, DC: American Psychiatric Publishing (2013).
- Fink P, Toft T, Hansen MS, Ørnbøl E, Olesen F. Symptoms and syndromes of bodily distress: an exploratory study of 978 internal medical, neurological, and primary care patients. *Psychosom Med.* (2007) 69:30–9. doi: 10.1097/PSY.0b013e31802e46eb
- Zijlema WL, Stolk RP, Löwe B, Rief W, White PD, Rosmalen JG. How to assess common somatic symptoms in large-scale studies: a systematic review of questionnaires. J Psychosom Res (2013) 74:459–68. doi: 10.1016/j.jpsychores.2013.03.093
- Witthöft M, Hiller W, Loch N, Jasper F. The latent structure of medically unexplained symptoms and its relation to functional somatic syndromes. *Int J Behav Med.* (2013) 20:172–83. doi: 10.1007/s12529-012-9237-2

- 22. Witthöft M, Fischer S, Jasper F, Rist F, Nater UM. Clarifying the latent structure and correlates of somatic symptom distress: a bifactor model approach. *Psychol Assess* (2016) 28:109. doi: 10.1037/pas0000150
- He X, Li C, Qian J, Cui H, Wu W. Reliability and validity of a generalized anxiety disorder scale in general hospital outpatients. *Shanghai Arch Psychiatry* (2010) 22:200–3. doi: 10.3969/j.issn.1002-0829.2010.04.002
- 24. Liu L, Bi B, Qin X, Wei S, Wang W, Li Y, et al. The prevalence of somatoform disorders in internal medicine outpatient departments of 23 general hospitals in Shenyang, China. *Gen Hosp Psychiatry* (2012) 34:339–44. doi: 10.1016/j.genhosppsych.2012.02.002
- 25. Kirmayer LJ, Sartorius N. Cultural models and somatic syndromes. *Psychosom Med.* (2007) 69:832–40. doi: 10.1097/PSY.0b013e31815b002c
- Escobar JI, Gureje O. Influence of cultural and social factors on the epidemiology of idiopathic somatic complaints and syndromes. *Psychosom Med.* (2007) 69:841–5. doi: 10.1097/PSY.0b013e31815b007e
- Hinton DE, Kredlow MA, Pich V, Bui E, Hofmann SG. The relationship of PTSD to key somatic complaints and cultural syndromes among Cambodian refugees attending a psychiatric clinic: The Cambodian Somatic Symptom and Syndrome Inventory (CSSI). *Transcult Psychiatry* (2013) 50:347–70. doi: 10.1177/1363461513481187
- Hong GK, Lee BS, Lorenzo MK. Somatization in Chinese American clients: Implications for psychotherapeutic services. J Contemp Psychother. (1995) 25:105–18. doi: 10.1007/BF02306684
- 29. Lee S. Estranged bodies, simulated harmony, and misplaced cultures: neurasthenia in contemporary Chinese society. *Psychosom Med.* (1998) 60:448–57. doi: 10.1097/00006842-199807000-00010
- Lin K-M, Cheung F. Mental health issues for Asian Americans. *Psychiatric Serv.* (1999) 50:774–80. doi: 10.1176/ps.50.6.774
- Lee S, Yu H, Wing Y, Chan C, Lee AM, Lee DT, et al. Psychiatric morbidity and illness experience of primary care patients with chronic fatigue in Hong Kong. Am J Psychiatry (2000) 157:380–4. doi: 10.1176/appi.ajp.157. 3.380
- AlHadi AN, AlAteeq DA, Al-Sharif E, Bawazeer HM, Alanazi H, AlShomrani AT, et al. An arabic translation, reliability, and validation of Patient Health Questionnaire in a Saudi sample. *Ann Gen Psychiatry* (2017) 16:32. doi: 10.1186/s12991-017-0155-1
- 33. Montalban SR, Vives AC, Garcia-Garcia M. Validation of the Spanish version of the PHQ-15 questionnaire for the evaluation of physical symptoms in patients with depression and/or anxiety disorders: DEPRE-SOMA study. Actas Esp Psiquiatr. (2010) 38:345–57. Available online at: http://www.actasespanolasdepsiquiatria.es/repositorio/11/68/ENG/11-68-ENG-345-357-416443.pdf
- Han C, Pae C-U, Patkar AA, Masand PS, Kim KW, Joe S-H, et al. Psychometric properties of the Patient Health Questionnaire-15 (PHQ-15) for measuring the somatic symptoms of psychiatric outpatients. *Psychosomatics* (2009) 50:580-5. doi: 10.1176/appi.psy.50.6.580
- Lipowski ZJ. Somatization: the concept and its clinical application. Am J Psychiatry (1988) 145:1358. doi: 10.1176/ajp.145.11.1358
- Van der Feltz-Cornelis C, Van Dyck R. The notion of somatization: An artefact of the conceptualization of body and mind. *Psychother Psychosom.* (1997) 66:117–27. doi: 10.1159/000289121
- Van der Feltz-Cornelis CM, van Balkom AJ. The concept of comorbidity in somatoform disorder - a DSM-V alternative for the DSM-IV classification of somatoform disorder. J Psychosom Res. (2010) 68:97–9. doi: 10.1016/j.jpsychores.2009.09.011
- Dere J, Sun J, Zhao Y, Persson TJ, Zhu X, Yao S, et al. Beyond "somatization" and "psychologization": symptomlevel variation in depressed Han Chinese and Euro-Canadian outpatients. *Front Psychol.* (2013) 4:377. doi: 10.3389/fpsyg.2013. 00377
- Xiong N, Wei J, Fritzsche K, Leonhart R, Hong X, Li T, et al. Psychological and somatic distress in Chinese outpatients at general hospitals: a cross-sectional study. Ann Gen Psychiatry (2017) 16:35. doi: 10.1186/s12991-017-0158-y
- 40. de Vroege L, Timmermans A, Kop WJ, van der Feltz-Cornelis CM. Neurocognitive dysfunctioning and the impact of comorbid depression and anxiety in patients with somatic symptom and related disorders: a cross-sectional clinical study. *Psychol Med.* (2017) 4:1–11. doi: 10.1017/S0033291717003300

- 41. van Meerkerk-Aanen PJ, de Vroege L, Khasho D, Foruz A, van Asseldonk JT, van der Feltz-Cornelis CM. La belle indifférence revisited: a case report on progressive supranuclear palsy misdiagnosed as conversion disorder. *Neuropsychiatr Dis Treat.* (2017) 13:2057. doi: 10.2147/NDT.S130475
- 42. van der Feltz-Cornelis CM, Andrea H, Kessels E, Duivenvoorden HJ, Biemans H, Metz M. Shared decision making in combinatie met ROM bij patiënten met gecombineerde lichamelijke en psychische klachten; een klinisch-emperische verkenning [article in Dutch] English: does routine outcome monitoring have a promising future? An investigation into the use of shred decision-making combined with ROM for patients with a combination of physical and psychiatric symptoms. *Tijdschrift Psychiatr.* (2014) 56:375–84. Available online at: http://www.tijdschriftvoorpsychiatrie.nl/issues/478/articles/10323
- Spitzer RL, Kroenke K, Williams JB, Group PHQPCS. Validation and utility of a self-report version of PRIME-MD: the PHQ primary care study. *JAMA* (1999) 282:1737–44. doi: 10.1001/jama.282.18.1737
- 44. Gierk B, Kohlmann S, Toussaint A, Wahl I, Brünahl CA, Murray AM, et al. Assessing somatic symptom burden: a psychometric comparison of the Patient Health Questionnaire--15 (PHQ-15) and the Somatic Symptom Scale--8 (SSS-8). J Psychosom Res. (2015) 78:352-5. doi: 10.1016/j.jpsychores.2014.11.006
- Kocalevent R-D, Hinz A, Brähler E. Standardization of a screening instrument (PHQ-15) for somatization syndromes in the general population. *BMC Psychiatry* (2013) 13:91. doi: 10.1186/1471-244X-13-91
- Lee S, Ma YL, Tsang A. Psychometric properties of the Chinese 15-item Patient Health Questionnaire in the general population of Hong Kong. J Psychosom Res. (2011) 71:69–73. doi: 10.1016/j.jpsychores.2011.01.016
- 47. Qian J, Ren Z, Yu D, He X, Li C. The value of the Patient Health Questionnaire-15 (PHQ-15) for screening somatic symptoms in general hospital. *Chin Ment Health J.* (2014) 28:173–8. Available online at: https:// caod.oriprobe.com/articles/41291902/The\_value\_of\_the\_Patient\_Health\_ Questionnaire\_15\_\_PHQ\_15\_for\_screenin.htm
- Muntingh A, Van Der Feltz-cornelis C, Van Marwijk H, Spinhoven P, Assendelft W, De Waal M, et al. Effectiveness of collaborative stepped care for anxiety disorders in primary care: a pragmatic cluster randomised controlled trial. *Psychother Psychosom.* (2014) 83:37–44. doi: 10.1159/0003 53682
- 49. Huijbregts KM, De Jong FJ, Van Marwijk HW, Beekman AT, Adèr HJ, Hakkaart-van Roijen L, et al. A target-driven collaborative care model for Major Depressive Disorder is effective in primary care in the Netherlands. A randomized clinical trial from the depression initiative. *J Affect Disord.* (2013) 146:328–37. doi: 10.1016/j.jad.2012.09.015
- Volker D, Zijlstra-Vlasveld MC, Anema JR, Beekman AT, Brouwers EP, Emons WH, et al. Effectiveness of a blended web-based intervention on return to work for sick-listed employees with common mental disorders: results of a cluster randomized controlled trial. *J Med Internet Res.* (2015) 17:e116. doi: 10.2196/jmir.4097
- 51. van Steenbergen-Weijenburg KM, van der Feltz-Cornelis CM, van Benthem TB, Horn EK, Ploeger R, Brals JW, et al. Collaborative care for co-morbid major depressive disorder in chronically ill outpatients in a general hospital. *Tijdschrift Psychiatr.* (2015) 57:248–57. Available online at: http://www. tijdschriftvoorpsychiatrie.nl/en/issues/488/articles/10543
- van Ravesteijn H, Wittkampf K, Lucassen P, van de Lisdonk E, van den Hoogen H, van Weert H, et al. Detecting somatoform disorders in primary care with the PHQ-15. *Ann Fam Med.* (2009) 7:232–8. doi: 10.1370/af m.985
- de Vroege L, Hoedeman R, Nuyen J, Sijtsma K, van der Feltz-Cornelis CM. Validation of the PHQ-15 for somatoform disorder in the occupational health care setting. *J Occup Rehabil.* (2012) 22:51–8. doi: 10.1007/s10926-011-9 320-6
- Löwe B, Gräfe K, Zipfel S, Witte S, Loerch B, Herzog W. Diagnosing ICD-10 depressive episodes: superior criterion validity of the Patient Health Questionnaire. *Psychother Psychosom.* (2004) 73:386–90. doi: 10.1159/000080393

- Löwe B, Kroenke K, Herzog W, Gräfe K. Measuring depression outcome with a brief self-report instrument: sensitivity to change of the Patient Health Questionnaire (PHQ-9). J Affect Disord. (2004) 81:61–6. doi: 10.1016/S0165-0327(03)00198-8
- Lowe B, Unutzer J, Callahan CM, Perkins AJ, Kroenke K. Monitoring depression treatment outcomes with the patient health quessionaire-9. *Medical Care* (2004) 42:1194–201. doi: 10.1097/00005650-200412000-00006
- Chen S, Chiu H, Xu B, Ma Y, Jin T, Wu M, et al. Reliability and validity of the PHQ-9 for screening late-life depression in Chinese primary care. *Int J Geriatr. Psychiatry* (2010) 25:1127–33. doi: 10.1002/gps.2442
- Spitzer RL, Kroenke K, Williams JB, Löwe B. A brief measure for assessing generalized anxiety disorder: the GAD-7. *Arch Intern Med.* (2006) 166:1092–7. doi: 10.1001/archinte.166.10.1092
- Löwe B, Decker O, Müller S, Brähler E, Schellberg D, Herzog W, et al. Validation and standardization of the Generalized Anxiety Disorder Screener (GAD-7) in the general population. *Med Care* (2008) 46:266–74. doi: 10.1097/MLR.0b013e318160d093
- 60. Muntingh AD, De Heer EW, Van Marwijk HW, Adèr HJ, Van Balkom AJ, Spinhoven P, et al. Screening high-risk patients and assisting in diagnosing anxiety in primary care: the Patient Health Questionnaire evaluated. BMC Psychiatry (2013) 13:192. doi: 10.1186/1471-244X-13-192
- 61. IBM Statistics. SPSS Version 24.0. Chicago, IL (2016).
- 62. Muthén LK, Muthén BO. Mplus. Statistical Analysis with Latent Variables Version. Los Angeles, CA (2007).
- Hu Lt, Bentler PM. Cutoff criteria for fit indexes in covariance structure analysis: Conventional criteria versus new alternatives. *Struct Equat Model Multidiscipl J.* (1999) 6:1–55.
- Kleinman A. Neurasthenia and depression: a study of somatization and culture in China. Cult Med Psychiatry (1982) 6:117–90. doi: 10.1007/BF00051427
- Ryder AG, Chentsova-Dutton YE. Depression in cultural context: "Chinese somatization," revisited. *Psychiatr Clin N Am.* (2012) 35:15–36. doi: 10.1016/j.psc.2011.11.006
- Ryder AG, Ban LM, Chentsova-Dutton YE. Towards a culturalclinical psychology. Soc Pers Psychol Compass (2011) 5:960–75. doi: 10.1111/j.1751-9004.2011.00404.x
- 67. Kleinman A, Kleinman J. Remembering the cultural revolution: alienating pains and the pain of alienation/transformation. In: Lin TY, Tsen W-S, Yeh E editors, *Mental Health in Chinese Societies*, Hong Kong: Oxford University Press (1995), pp. 141–55.
- Kleinman A, Anderson J, Finkler K, Frankenberg R, Young A. Social origins of distress and disease: depression, neurasthenia, and pain in modern China. *Curr Anthropol.* (1986) 24:499–509.
- 69. Keen N, George D, Scragg P, Peters E. The role of shame in people with a diagnosis of schizophrenia. *Brit J Clin Psychol.* (2017) 56:115–29. doi: 10.1111/bjc.12125
- 70. Xu X, Li X-M, Zhang J, Wang W. Mental health-related stigma in China. Issues Ment Health Nurs. (2017) 39:126–134. doi: 10.1080/01612840.2017.1368749
- Zhou X, Dere J, Zhu X, Yao S, Chentsova-Dutton YE, Ryder AG. Anxiety symptom presentations in Han Chinese and Euro-Canadian outpatients: is distress always somatized in China? *J Affect Disord*. (2011) 135:111–4.

**Conflict of Interest Statement:** The authors declare that the research was conducted in the absence of any commercial or financial relationships that could be construed as a potential conflict of interest.

Copyright © 2018 Leonhart, de Vroege, Zhang, Liu, Dong, Schaefert, Nolte, Fischer, Fritzsche and van der Feltz-Cornelis. This is an open-access article distributed under the terms of the Creative Commons Attribution License (CC BY). The use, distribution or reproduction in other forums is permitted, provided the original author(s) and the copyright owner are credited and that the original publication in this journal is cited, in accordance with accepted academic practice. No use, distribution or reproduction is permitted which does not comply with these terms.