REHABILITATION & REGENERATIVE MEDICINE SECTION

Which Chronic Low Back Pain Patients Respond Favorably to Yoga, Physical Therapy, and a Self-care Book? Responder Analyses from a Randomized Controlled Trial

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Abstract

Purpose. To identify baseline characteristics of adults with chronic low back pain (cLBP) that predict response (i.e., a clinically important improvement) and/or modify treatment effect across three nonpharmacologic interventions. Design. Secondary analysis of a randomized controlled trial. Setting. Academic safety net hospital and seven federally qualified community health centers. Subjects. Adults with cLBP (N = 299). Methods. We report patient characteristics that were predictors of response and/or modified treatment effect across three 12-week treatments: yoga, physical therapy [PT], and a self-care book. Using preselected characteristics, we used logistic regression to identify predictors of "response," defined as a >30% improvement in the Roland Morris Disability Questionnaire. Then, using "response" as our outcome, we identified baseline characteristics that were treatment effect modifiers by testing for statistical interaction (P < 0.05) across two comparisons: 1) yoga-or-PT vs self-care and 2) yoga vs PT. Results. Overall, 39% (116/299) of participants were responders, with more responders in the yoga-or-PT group (42%) than the self-care (23%) group. There was no difference in proportion responding to voga (48%) vs PT (37%, odds ratio [OR] = 1.5, 95% confidence interval = 0.88 - 2.6). Predictors of response included having more than a high school education, a higher income, employment, few depressive symptoms, lower perceived stress, few work-related fear avoidance beliefs, high pain self-efficacy, and being a nonsmoker. Effect modifiers included use of pain medication and fear avoidance beliefs related to physical activity (both P=0.02 for interaction). When comparing yoga or PT with self-care, a greater proportion were responders among those using pain meds (OR = 5.3), which differed from those not taking pain meds (OR = 0.94) at baseline. We also found greater treatment response among those with lower (OR = 7.0), but not high (OR = 1.3), fear avoidance beliefs around physical activity. **Conclusions**. Our findings revealed important subgroups for whom referral to yoga or PT may improve cLBP outcomes.

Key Words: Chronic Low Back Pain; Yoga; Physical Therapy; Primary Care; Predictor of Response; Effect Modification; Health Disparities

Introduction

Recent clinical practice guidelines from the American College of Physicians recommend multiple nonpharmacologic treatments as safe and effective first-line therapy for chronic low back pain (cLBP) [1]. However, patients may respond differently to treatments [2]. Determining who benefits from what treatment will help guide care. This information may be particularly useful in primary care settings where patients and providers are potentially overwhelmed by a long list of similarly effective treatment options [3, 4].

Yoga, a mind and body practice that includes postures, breathing, meditation, and relaxation, is increasingly popular in the United States and is often practiced in groups [5]. The most common nonpharmacologic referral from primary care physicians for adults with cLBP is to physical therapy (PT), where physical therapists tailor treatment to individuals during one-on-one sessions [6]. Aerobic exercise and strength training are important and common components of PT. While systematic reviews suggest that yoga [7, 8] and PT [9] improve back-related pain and physical function, there is little guidance on which approach may be most effective for a particular patient.

One focus of this paper is to identify those patient characteristics that predict a clinically meaningful improvement in cLBP, which we call "predictors of response." This is in part based on the National Institutes of Health (NIH) Task Force on Research Standards for Chronic Low Back Pain, which recommended the use of responder analyses to identify individuals who have a clinically meaningful improvement, for example, a 30% improvement in back-related physical function [10-12]. Baseline characteristics can be used to predict negative or positive clinical outcomes [13]. For example, in studies of PT, predictors of negative outcomes (e.g., chronicity, no improvement) include referred leg pain, catastrophizing, fear, anxiety, and depression [14]. Similarly, predictors of positive outcomes have been identified in studies of yoga [15, 16] and particular PT treatments, for example, flexion/extension exercises [17–19]. This implies that some factors may generally predict favorable outcomes, although this information may not help clinicians choose between treatments.

It may be more interesting and useful to understand whether characteristics can define subgroups that may respond differently to particular treatments, which we term "effect modification." Thus, a second focus of this paper is to identify treatment effect modifiers, which are baseline characteristics that influence outcomes by interacting with the treatment received. Characteristics that are

treatment effect modifiers can be used to delineate subgroups of patients that are more or less likely to achieve a 30% improvement (i.e., response) in physical function across different interventions (e.g., yoga, PT, and selfcare) [13]. Identifying subgroups that guide treatment is a longstanding priority for low back pain research agendas [10, 20]. Such subgroups can help inform common questions in primary care settings: Which patients will do well with self-care only? Who may need more structured or supervised treatment? Who will do better with a particular treatment (e.g., yoga or PT)? Yet, treatment effect modification analyses are challenging, as they require measurement of baseline factors of interest and a large sample size. Thus, few clinical trials have evaluated effect modifiers of cLBP treatments [21, 22], and information on effect modifiers from clinical trials comparing similarly effective nonpharmacologic treatments is particularly scant.

The Back to Health randomized controlled trial, which recruited participants with cLBP from low-income and racially diverse communities, found yoga to be non-inferior to PT for improving back-specific physical function and reducing pain [23, 24]. The purpose of these secondary analyses was to address two additional important clinical questions: First, what pretreatment characteristics are important predictors of response in this understudied and underserved population? Second, what pretreatment characteristics are effect modifiers in the following two comparisons: 1) yoga or PT compared with a self-care book and 2) yoga vs PT.

Methods

Study Design

This is a secondary analysis of a three-arm clinical trial where adults with cLBP were randomized to yoga, PT, or self-care in a 2:2:1 ratio [23, 24]. The sample size for the original study (N = 320) was determined by power calculations for the primary aim, which tested whether yoga was noninferior to physical therapy on pain and backrelated physical function outcomes. The methods [23] and primary results [24] are described in detail elsewhere and summarized briefly below. Participants were recruited from predominantly low-income racially diverse neighborhoods in Boston, Massachusetts. This study was approved by the Boston University Medical Campus Institutional Review Board. All participants provided written informed consent.

Participants

The complete set of inclusion and exclusion criteria were reported previously [23]. Eligible participants were

English-speaking adults (ages 18–64) who had low back pain persisting \geq 12 weeks with an average pain intensity in the previous week \geq 4 on an 11-point numerical rating scale. Individuals with suspected or confirmed specific causes of back pain were excluded, for example, spinal stenosis, radiculopathy, or those who had red flags indicating serious pathology (e.g., cancer, infection). Individuals were also excluded if they had an active or planned workers' compensation, disability, or personal injury claim. For this secondary analysis, we additionally excluded 21 participants with missing data at 12 weeks on our main outcome, the Roland Morris Disability Questionnaire.

Study Interventions

Our manualized hatha yoga intervention consisted of 12 group-based weekly 75-minute classes incorporating poses, relaxation/meditation exercises, yoga breathing, and yoga philosophy [23–26]. Thirty minutes of daily home practice was encouraged and supported with supplies (yoga mat, blocks, instructional DVD/manual).

Our manualized PT intervention consisted of 15 oneon-one 60-minute appointments over 12 weeks [23, 24]. During each appointment, the physical therapist utilized the Treatment-Based Classification Method and supervised aerobic exercise [27, 28]. Participants with a high score on the Fear Avoidance Beliefs Questionnaire (FABQ) work subscale (>29) received *The Back Book* [29], a brief educational resource. Physical therapists reinforced its principles to lower fear avoidance [29]. Physical therapists provided written instructions and supplies for exercises to be practiced daily at home [23]. Participants were encouraged by physical therapists to log these prescribed exercises and the number of repetitions completed.

Participants receiving the self-care intervention were provided a copy of *The Back Pain Helpbook* [30], a comprehensive resource describing evidence-based self-management strategies for cLBP, including stretching, strengthening, and the role of psychological and social factors. Every three weeks, participants received a oneto two-page newsletter summarizing assigned chapters and a five- to 10-minute check-in telephone call from staff.

Data Collection

For this exploratory analysis, we identified patient-level characteristics from multiple domains (sociodemographic, general health, back-related, psychological, and treatment expectations) [31]. Data were collected with surveys administered by study staff blinded to treatment. We restricted our investigation to characteristics previously identified, or clinically plausible, as predictors of response or effect modifiers [13, 21, 22]. Characteristics were measured before randomization with valid and reliable instruments [13, 23].

Sociodemographic Characteristics

Baseline sociodemographic characteristics were age (18– 44, 45–65), sex, race (white, nonwhite), ethnicity (Hispanic, non-Hispanic), household income (\leq \$30,000, >\$30,000), education (high school or less, more than high school), and current employment (employed, unemployed).

General Health Measures

General health characteristics were obesity (body mass index $[BMI] > 30 \text{ kg/m}^2$), current smoker, and any exercise in the previous week. We characterized participants as having multiple chronic comorbidities if they self-reported two or more of the following conditions: hypertension, osteoarthritis, neck pain, fibromyalgia, depression, other psychiatric disorder, diabetes, or chronic obstructive pulmonary disease.

Back-Related Measures

We used the 11-point numeric rating scale (0-10) to identify participants with moderate (4-6) or severe (≥ 7) back pain [32, 33]. We defined medication use as any use of one or more medications in the following categories in the previous week: nonsteroidal anti-inflammatory drugs (NSAIDs), acetaminophen, muscle relaxants, or an opioid medication. Participants reported the number of days of any back pain in the previous three months and activity-limiting back pain in the past month. Each of these measures was dichotomized at the median number of days.

Psychological Measures

Baseline psychological factors included symptoms of depression (≥ 10 on PHQ-8 [34]) and anxiety (≥ 10 on the Generalized Anxiety Disorder 7-item questionnaire [GAD-7] [35]). Perceived Stress Scale values were dichotomized using the median score to identify those with elevated perceived stress (≥ 17 on PSS-10) [36]. Fear avoidance relating to physical activity (>13 on FABQ-PA) and work (>29 on FABQ-W) was identified [37]. Using the Pittsburgh Sleep Quality Index (PSQI) [38], we identified participants with poor sleep (score >5). We classified participants as having maladaptive pain coping skills, using a score ≥ 14 on the catastrophizing subscale of the Coping Strategies Questionnaire (CSQ) [39], and low to moderate pain self-efficacy, using a score <40 on the Pain Self-Efficacy Questionnaire (PSEQ) [40, 41].

Treatment Expectations

Participants were asked before randomization whether they expected each of the treatment arms would be helpful for their cLBP using an 11-point scale. Those who responded 9 or 10 were considered to have high expectations. We identified concordance in expectations with treatment allocation, for example, someone who expected PT would be helpful and was then randomized to PT.

Definition of a Clinically Important Response

The 23-item modified Roland Morris Disability Questionnaire (RMDQ) is a validated, widely used measure of back-specific physical function, where higher values indicate worse function [42]. Participant response was calculated as the percent change in Roland Morris score from baseline to 12-week follow-up. We defined "response" as achieving a minimal clinically important difference on the RMDQ (\geq 30% improvement) [11, 12]. The reference group in all analyses was individuals not meeting this response threshold.

Data Analysis

We compared the proportion of responders by baseline characteristics using chi-square and *t* tests for categorical and continuous variables, respectively. Possible predictors of "response" were selected based on a nominal univariate *P* value of <0.1. Odds ratios with 95% confidence intervals were calculated for each selected predictor after adjusting for age, baseline RMDQ, and treatment group.

To assess for effect modification, we developed separate logistic regression models for each baseline characteristic to predict "response" as the outcome. Models were adjusted for baseline age and RMDQ and included an interaction term (characteristic*treatment). Each model adjusted for age and baseline RMDQ. We assessed for effect modification in the two following comparisons: 1) yoga or PT vs self-care book or 2) yoga vs PT. We chose these comparisons as they reflect questions patients often ask their primary care providers and have the potential to inform shared decision-making (as illustrated in Appendix Figure 1). For the two comparisons, we generated separate predictor-stratified estimates (odds ratios) of response for each baseline characteristic assessed. To account for zero responders in certain subgroups, where estimates could not be obtained with a standard logistic regression model, we employed Firth's penalty to estimate the odds ratio and approximate confidence bounds [43, 44]. A statistically significant interaction term (P < 0.05) indicated effect modification. A P value of 0.05 to 0.20 indicated exploratory evidence for effect modification [21, 22].

SAS, version 9.4 (SAS Institute, Cary, NC, USA), was used for all analyses.

Results

Sample Characteristics

Among 299 participants with follow-up data, most were nonwhite (82%), female (66%), and had an annual income of \$30,000 or less (64%) (Appendix Table 1). Roughly half were currently employed (45%) at the beginning of the study. The characteristics of those with complete data on RMDQ at 12 weeks (N=299) were generally similar to those with missing outcome data (N=21) on sociodemographic and health characteristics, although participants with complete data were more likely to be female than those with missing data (66% vs 38%, P = 0.01) (Appendix Table 2).

Treatment Characteristics

Details on attendance of sessions and home practice participation have been reported previously [24]. Briefly, the median attendance was seven visits and seven sessions for yoga and PT participants, respectively. During the 12week intervention phase, yoga participants reported practicing at home four days per week (median = 27 minutes). Similarly, PT participants reported practicing home exercises four days per week (median = 4 exercises).

Predictors of a Clinically Important Response

Thirty-nine percent (116/299) of all study participants experienced a clinically meaningful improvement in back-specific physical function at 12 weeks (Figure 1). Predictors of response included being a nonsmoker (vs current smoker, 47% vs 21%, P < 0.0001), not using pain medication (vs using any, 54% vs 32%, P < 0.0001), being employed (vs unemployed, 52% vs 28%, P < 0.0001), income >\$30,000 (vs \leq \$30,000, 54% vs 31%, P = 0.0002), low/moderate depressive symptoms on the PHQ-8 (vs high, 48% vs 23%, P = 0.0005), lower perceived stress on the PSS-10 (vs higher, 46% vs 32%, P = 0.004), fewer work-related fear avoidance beliefs on the FABQ-W (vs high, 43% vs 17%, P = 0.001), high pain self-efficacy on the PSEQ (vs low/ moderate, 49% vs 31%, P = 0.001), and receiving treatment they expected would be helpful, that is, concordance (vs not, 47% vs 33%, P = 0.02) (Appendix Table 3).

Table 1 shows odds ratios for important factors adjusted for age, baseline RMDQ, and treatment group. These variables are presented out of all the a priori risk factors based on a univariate P value <0.1, and due to having the largest absolute value standardized beta estimates in a multivariate model that included all potential predictors.

Treatment Effect Modifiers

In Figure 2, we provide four illustrative examples of baseline characteristics that are neither a predictor of response nor a treatment effect modifier (Figure 2a), that are a predictor of response only (Figure 2b), that are an effect modifier only (Figure 2c), and that are both a predictor of response and an effect modifier (Figure 2d). Table 2 presents the results of separate adjusted regression models to determine treatment effect modification in the following two comparisons.



Figure 1. Flow diagram of postintervention response comparisons for yoga, physical therapy, and self-care.

Yoga or PT vs Self-care

Participants receiving yoga or PT were more likely to be responders than participants receiving self-care (43% vs 23%, OR = 2.7, 95% CI = 1.4 - 5.2). Two baseline characteristics showed clear evidence of treatment effect modification: pain medication use (P = 0.02) and FABQ-PA (P = 0.02) (Figure 2d and c, respectively). Participants (N = 210) using pain medications at baseline were 5.3 times more likely to be responders at week 12 if they were in the yoga or PT group compared with self-care (38% vs 11%, OR = 5.3, 95% CI = 2.0 - 14.2). In contrast, among 89 participants not taking pain medications, the proportion of responders was not statistically different in yoga or PT compared with self-care (53% vs 56%, OR = 0.94, 95% CI = 0.31 - 2.9). This pattern was consistent for use of individual medication categories (acetaminophen, NSAIDs, and opioids), as shown in Appendix Table 3. Similarly, while 178 participants with low/moderate FABQ-PA scores were more likely to be responders with yoga or PT compared with self-care (47% vs 13%, OR = 7.0, 95% CI = 2.2 - 21.8), this was not observed for 121 participants with initial high FABQ-PA scores (38% vs 31%, OR = 1.3, 95% CI = 0.56 - 3.2). These findings, that pain medication use and

fear avoidance beliefs around physical activity were effect modifiers, were also observed after additionally adjusting for baseline pain intensity.

Characteristics with exploratory evidence for effect modification included educational attainment (P = 0.14), patient expectations for yoga (P = 0.14), and having expectations concordant with treatment allocation (P = 0.06). Firth's penalty was used to estimate odds ratios for strata of two characteristics (i.e., educational attainment, smoking) where there were zero responders in one of the treatment groups (Table 2). For example, among participants with less than a high school education randomized to self-care, zero were responders, compared with a third of those receiving yoga or PT.

Yoga vs PT

Yoga was similarly effective compared with PT (48% vs 37%, OR = 1.5, 95% CI = 0.88 – 2.6). Among 121 participants who expected yoga would be helpful, more benefited from yoga vs PT (63% vs 39%, OR = 2.5, 95% CI = 1.1–5.8). Among those who had lower expectations for yoga, there were similar proportions of responders for yoga and PT (38% vs 36%, OR = 1.0,

Baseline Characteristics	Responders, No. (%)	Adjusted, * OR (95% CI)
Age per year aged		0.97 (0.95, 1.00)
Reland Morris	-	0.97(0.93-1.00)
nor 1 point increase	-	0.97 (0.92-1.02)
Education		
Lish school on loss	22/11/ (28)	1.0
Mana than high ashaal	52/116 (28) 81/180 (45)	1.0
More than high school	81/180 (45)	2.1 (1.2-3.3)
	54/175 (21)	0 45 (0 26 0 70)
≤\$30,000 ≥ \$20,000	52/00 (51)	0.43 (0.26-0.78)
>\$30,000	33/99 (34)	1.0
Linployment N-	46/164 (29)	1.0
NO Yaa	46/164 (28)	1.0
Tes Comochid conditions	/0/135 (52)	2.5 (1.5–4.2)
	70/150/50	1.0
0-1	/9/158 (50)	1.0
≥ 2	3//141 (26)	0.43 (0.26-0.71)
Current smoker	07/200 (47)	1.0
No	9//208 (4/)	1.0
Yes	19/91 (21)	0.31 (0.1/-0.5/)
Pain medication use	10/00 (54)	1.0
None	48/89 (54)	1.0
Any Dia	68/210 (32)	0.46 (0.2/-0./8)
Depression symptoms, PHQ-8	02/102 (40)	1.0
Low/moderate, 0–9	93/192 (48)	1.0
High, ≥ 10	23/102 (23)	0.28 (0.15-0.51)
Perceived stress, PSS-10	(11120 (10)	1.0
Low(<1/)	64/138 (46)	1.0
High (≥ 17)	50/157 (32)	0.51 (0.31-0.85)
Fear avoidance,		
FABQ physical activity		
Low/moderate, 0–13	56/141 (40)	1.0
High, >13	57/155 (37)	0.85 (0.52–1.4)
Fear avoidance, FABQ work		
Low/moderate, 0–29	105/246 (43)	1.0
High-risk, >29	7/42 (17)	0.32 (0.14–0.77)
Pain self-efficacy, PSEQ		
Low/moderate (<40)	51/166 (31)	0.46 (0.27–0.79)
High (≥ 40)	65/132 (49)	1.0
Expectations, concordant		
Received a less	60/180 (33)	1.0
"helpful" treatment		
Received a "helpful" treatment	56/119 (47)	1.5 (0.94–2.5)
Treatment group		
Yoga	60/125 (48)	3.3 (1.6-6.7)
РТ	42/113 (37)	2.2 (1.1-4.5)
Self-care	14/60 (23)	1.0
Treatment group		
comparisons of interest		
Yoga or PT vs self-care	-	2.7 (1.4-5.2)
Yoga vs PT	-	1.5 (0.88-2.6)

 Table 1. Predictors of 30% improvement in back-specific physical function on the Roland Morris Disability Questionnaire among 299 participants

CI = confidence interval; FABQ = Fear Avoidance Beliefs Questionnaire; OR = odds ratio; PHQ-8 = Patient Health Questionnaire; PSEQ = Pain Self-Efficacy Questionnaire; PSS-10 = 10-item Perceived Stress Scale.

*Adjusted for baseline age, baseline Roland Morris score, and treatment group.

95% CI = 0.51–2.1). However, this finding did not meet the criteria for effect modification (P = 0.11). Additionally, this trend was not found among the 141 participants who expected to do well with PT (Table 2).

No other potential effect modifiers for the yoga vs PT comparison were identified. Moreover, factors identified as effect modifiers of the yoga or PT vs self-care comparison were not effect modifiers of the yoga vs PT comparison, that is, pain medication use (P = 0.54) and FABQ-PA (P = 0.87) (Figure 3).

Discussion

In a predominantly nonwhite low-income sample of 299 adults with cLBP, predictors of response independent of treatment included more than a high school education, higher income, employment, zero or one comorbid chronic conditions, not smoking, not using pain medication, low depressive symptoms, low perceived stress, few work-related fear avoidance beliefs, and higher pain selfefficacy. Participants who received either yoga or PT were more likely to achieve clinically important improvements in physical function compared with self-care. Although this treatment effect was consistent across a range of baseline characteristics, the magnitude of responders in yoga or PT compared with self-care was augmented among participants already using pain medication and those with few fear avoidance beliefs related to physical activity. The only potential effect modifier between yoga and PT was treatment expectation: Participants who expected to do well with yoga did better with yoga than PT.

The average income of our population is well below the US median income, and our study adds important data for this understudied and often underserved population. We observed predictors of response consistent with previous literature; that is, lower socioeconomic status, multiple comorbidities, depression, and smoking are associated with poor response [45-48]. The majority of study participants were nonresponders at the end of the intervention period. Our interventions included predominantly physical and mind-body approaches. Participants with multiple comorbidities, depression, etc., may require a more intensive or comprehensive approach to experience a clinically meaningful improvement, that is, one that not only focuses on back pain, but also targeting behavioral modification and other physical or psychological comorbidities. Although we used only one discipline in each arm (physical therapist, yoga instructor), some evidence suggests that patients with more complex cLBP may benefit from multidisciplinary approaches [49, 50].

Our findings corroborate those of studies suggesting that medication use may modify the effect of treatments for low back pain [21, 22, 51] and other musculoskeletal conditions [52]. Among participants not using pain medications at baseline, the proportion of responders was relatively high for all three treatment groups. Among those using pain medication, we observed a large effect of yoga or PT compared with self-care. Current use of one or more pain medications may be a proxy for more persistent or otherwise severe cLBP where self-care may not be -

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		Responders, No. (%)						
Characteristic	Sample (N = 299)	Yoga (N = 125)	PT (N = 113)	S-C (N=61)	Yoga/PT vs S-C OR (95% CI)*	P Value†	Yoga vs PT OR (95% CI)*	P Value†
Education‡						0.14		0.80
<high school<="" td=""><td>116</td><td>19/49 (39)</td><td>13/49 (27)</td><td>0/18 (0)</td><td>19.5 (1.1-360.3)‡</td><td></td><td>1.6 (0.69-3.9)</td><td></td></high>	116	19/49 (39)	13/49 (27)	0/18 (0)	19.5 (1.1-360.3)‡		1.6 (0.69-3.9)	
> High school	180	39/74 (53)	28/63 (44)	14/43 (33)	2.0 (0.97-4.2)		1.4 (0.71-2.9)	
Annual income						0.55		0.78
<\$30,000	175	28/75 (37)	20/62 (32)	6/38 (16)	2.4 (0.82-7.0)		1.7 (0.68-4.2)	
>\$30,000	99	27/43 (63)	19/38 (50)	7/18 (39)	2.9 (1.1-7.5)		1.2 (0.58-2.5)	
Employment						0.39		0.64
No	135	22/66 (33)	19/66 (29)	5/32 (16)	2.4 (0.86-6.7)		1.2(0.57-2.5)	
Yes	164	38/59 (64)	23/47 (49)	929 (31)	3.3 (1.4-8.2)		2.0(0.87-4.4)	
Comorbid conditions		. ,	· · ·	· · /	()	0.21	· · · · · ·	0.51
0–1	1.58	42/72 (58)	26/58 (45)	11/28 (39)	1.8(0.76-4.2)		1.7(0.85 - 3.6)	
>2	141	18/53 (34)	16/55 (29)	3/33 (9)	4.8 (1.4–16.9)		1.2(0.53-2.8)	
Smoker at baseline±						0.25		0.69
No	208	50/89 (56)	33/76 (43)	14/43 (33)	2.2 (1.1-4.6)		1.6(0.85 - 3.0)	
Yes	91	10/36 (28)	9/37 (24)	0/18 (0)	$13.3(0.71-248.7)\pm$		1.3(0.43-3.6)	
Pain medication use					(/+	0.02		0.54
None	89	24/40 (60)	15/33 (45)	9/16 (56)	0.94(0.31-2.9)	0.02	1.9(0.75 - 5.0)	0.01
Any	210	36/85 (42)	27/80 (34)	5/45 (11)	5.3(2.0-14.2)		1.3(0.70-2.6)	
Depression symptoms	210	00,00 (12)	2//00 (01)	0,10 (11)	010 (210 1112)	0.36	110 (01/ 0 210)	0.99
Low/moderate	192	47/82 (57)	33/71 (46)	13/39 (33)	25(12-53)	0.00	1.6(0.85-3.2)	0.,,,
High	102	13/41(32)	9/41 (22)	1/20 (5)	7.0(0.87-56.1)		$1.0(0.03 \ 9.2)$ 1.7(0.61-4.5)	
Perceived stress	102	10, 11 (02)	>, ··· (<u>==</u>)	1,20 (0)	,10 (010) 0011)	0.94	117 (0101 110)	0.73
Low	138	33/60 (55)	23/50 (46)	8/28 (29)	28(11-70)	0.91	14(0.65-3.2)	0.75
High	157	26/63 (41)	18/62 (29)	6/32(19)	2.6(0.99-7.1)		1.7(0.82 - 3.2) 1.7(0.82 - 3.7)	
Fear avoidance PA	157	20/03 (11)	10/02 (2))	0,32 (17)	2.0 (0.55 7.1)	0.02	1.7 (0.02 0.77)	0.87
Low/moderate	155	29/55 (53)	23/55 (42)	4/31 (13)	70(22-218)	0.02	1.5(0.69-3.3)	0.07
High	141	30/69 (43)	18/57 (32)	9/29 (31)	1.3(0.56-3.2)		1.6(0.77-3.5)	
Fear avoidance work	111	56/65 (15)	10/37 (32)	<i>)</i> , <u>2</u>) (31)	1.5 (0.50 5.2)	0 44	1.0 (0.77 5.57	0.56
Low/moderate	246	58/110 (53)	34/85(40)	13/51 (25)	30(15-61)	0.11	1.7(0.92-3.0)	0.50
High	42	2/14 (14)	4/19 (21)	1/9(11)	1.4 (0.14 - 14.3)		0.8(0.12-5.0)	
Pain self-efficacy	12	2/11(11)	1/1/(21)	1/2 (11)	1.1 (0.11 11.5)	0.21	0.0 (0.12 5.0)	0.75
Low/moderate	166	27/70 (39)	17/64 (27)	7/32 (22)	19(07347)	0.21	18(08337)	0.75
High	132	33/55 (60)	17704(27) 25/48(52)	7/29 (24)	4.3(1.7-11.2)		1.0(0.05-3.7) 1.5(0.65-3.3)	
Expectations yoga	152	55/55 (00)	25740 (52)	//2/(24)	4.3 (1.7–11.2)	0.14	1.5 (0.05-5.5)	0.11
Low/moderate	170	20/72 (20)	25/69 (26)	10/26 (28)	1.7(0.75,4.0)	0.14	10(05121)	0.11
High	178	20/73 (30)	23/69(36) 17/44(39)	10/36(28)	1.7(0.73-4.0)		1.0(0.31-2.1)	
Furnestations DT	121	52/52 (62)	1//44 (37)	4/23 (10)	4.0 (1.0-14.0)	0.26	2.5 (1.1-5.8)	0.40
Landara danata	150	22/66/49	10/50 (21)	0/24(26)	10/092 45)	0.36	10/099/00	0.40
Low/moderate	138	32/66 (48) 28/59 (47)	18/38(31)	5/34 (26) 5/27 (10)	1.9(0.82-4.3)		1.9(0.88-4.0) 1.2(0.56, 2.5)	
riign	141	28/39 (47)	24/33 (44)	3/2/ (19)	5.6 (1.5-10.1)	0.21	1.2 (0.36-2.3)	0.25
Expectation, self-care+	222	50/07/52)	20/77 (20)	14/40 (20)	21/1042	0.21	1 0 (0 0 4 2 2)	0.55
Low/moderate	223	30/9/ (32) 10/27 (32)	271//(38)	14/49 (29)	2.1 (1.0-4.3)		1.8 (0.94 - 3.3)	
гign	/3	10/2/ (3/)	12/34 (33)	0/12(0)	13.0 (0.76-293.6)‡	0.07	0.97 (0.34-2.8)	0.22
Expectations, concordant‡	100	20/72 (20)	10/50 (21)	14/40 (20)	1 5 (0 70 2 0)	0.06	1 2 (0 50 2 5)	0.33
INO V	180	28//3 (38)	18/58 (31)	14/49 (29)	1.5(0.70-3.0)		1.2(0.38-2.6)	
1 @\$	119	32/32 (62)	24/33 (44)	0/12 (0)	26.4 (1.4–310.1)‡		2.1 (0.95-4.6)	

CI = confidence interval; PA = physical activity; PT = physical therapy intervention group; S-C = self-care intervention group.

*Adjusted for baseline age and Roland Morris.

[†]Test for multiplicative interaction (i.e., statistical effect modification) between baseline characteristics and treatment.

[‡]Calculated using logistic regression with Firth's penalty for zero cells.

sufficient. Adjusting for baseline pain intensity did not attenuate our findings, and, therefore, alternative explanations should be explored in future research. Early referral to a more structured treatment with a therapist or instructor may improve outcomes.

Fear avoidance beliefs are important predictors of response [53] and may be an important cLBP treatment effect modifier [54]. Participants with lower fear avoidance beliefs around physical activity were much more likely to be responders to yoga (53%) or PT (42%) than self-care (13%). In contrast, among participants who started out with high fear avoidance around physical activity, the proportions of responders to yoga (43%), PT (32%), and self-care (31%) were not statistically



Figure 2. Characteristics that are predictors of response and/or treatment effect modifiers, or neither, when comparing yoga or PT with self-care. For each panel, the baseline characteristic is shown as a potential predictor of response on the left and a potential treatment effect modifier on the right. The corresponding *P* values are from logistic regression models adjusted for age and baseline back-specific physical function on the Roland Morris Disability Questionnaire. In panel a, sex was neither a predictor of response (P = 0.46) nor a treatment effect modifier (*P* for interaction = 0.69). In panel b, the number of chronic conditions was a predictor of response (P < 0.001) but not a treatment effect modifier (*P* for interaction = 0.69). In panel c, higher fear avoidance beliefs around physical activity was not a predictor of response (P = 0.60) but was a treatment effect modifier (*P* for interaction = 0.02). In panel d, baseline pain medicine use was both a predictor of response (P < 0.001) and a treatment effect modifier (*P* for interaction = 0.02).



Figure 3. Forest plot of effect modification on the multiplicative scale. Odds ratios indicate likelihood of being a responder. For example, participants using pain medications at baseline were more likely to be responders if they were in the yoga or physical therapy (PT) group compared with self-care group (odds ratio [OR] = 5.3, 95% confidence interval [CI] = 2.0 - 14.2). In contrast, among those not taking pain medications at baseline, the proportion of responders was not statistically different in yoga or PT compared with self-care (OR = 0.94, 95% CI = 0.31 - 2.9).

different. While the decrease in response rate among yoga and PT in the high–fear avoidance strata may have been expected, our finding that more individuals with high fear avoidance in the self-care group responded compared with those with lower fear is difficult to interpret. The self-care book offers substantial content on exercise and addresses fear related to physical activity. While this may be useful, important, or new information for adults with high fear avoidance related to physical activity, individuals with low baseline fear may have found this content to be less useful. In contrast, the yoga instructor or physical therapist has the opportunity to individualize care, coach adults with cLBP to meet care goals, and teach the patient experientially that increasing their activity appropriately does not necessarily lead to more pain or disability. This encouragement may only be helpful if it overcomes fear avoidance related to physical activity and increases hours of back-related exercise and selfefficacy, which were important mediators of improvement in another study of yoga and stretching for cLBP [55].

A review of treatment effect modifiers in back pain clinical trials identified potential effect modifiers including age, employment, treatment expectations, and educational attainment [21]. Educational attainment has been identified as an effect modifier in a more recent study of cLBP treatment [22] and randomized trials evaluating interventions for osteoarthritis [52] and chronic pain [56]. In our study, when comparing yoga and PT with self-care, educational attainment was the only socioeconomic characteristic identified as a potential effect modifier but was not statistically significant (P for interaction = 0.14). While this finding should be interpreted with caution, it is notable that zero of 18 participants with high school education or less randomized to the self-care book responded. In contrast, roughly a third of the individuals with high school education or less receiving yoga or PT had a clinically meaningful response. Patients with less educational attainment may be unlikely to respond to book-based selfcare interventions. This is consistent with a large body of literature about education, health literacy, back pain, and other health outcomes [57, 58]. Having the support of a therapist or instructor may help overcome the barriers to improvement related to lower educational attainment.

Our finding of treatment expectation as a potential effect modifier is also consistent with previously identified effect modifiers of cLBP [21] and osteoarthritis [52] treatment. Paradoxically, among 22 participants who expected to do well with self-care and then were randomized to self-care, none were responders. Furthermore, it appeared that those who expected to do well with yoga were more likely to be responders, especially when they were randomized to yoga. While these findings require replication, it may be reasonable for primary care providers to encourage patients with a strong interest in yoga to pursue it as part of their cLBP care [1].

The principal limitation of this study is a relatively small sample size to investigate back pain subgroups. However, our approach is aligned with recommendations for effect modification analyses using clinical trial data; that is, we evaluated characteristics that were clinically plausible as effect modifiers and were measured with valid instruments before randomization [13, 59]. Despite relatively limited statistical power, we identified statistically significant treatment effect modifiers using the nominal Pvalue of 0.05. As our analyses were exploratory in nature, we did not adjust for multiple comparisons [60]. Importantly, the characteristics demonstrating effect modification (pain medication, fear avoidance) and those trending toward effect modification (e.g., educational attainment, patient expectations) in our study are consistent with previous literature [21, 22, 52, 56]. Nonetheless, our findings need to be validated in a prospective study.

As this was a secondary analysis, we were limited to the information that was collected for the original study. Measures that further characterize cLBP (e.g., duration of pain) and other patient characteristics (e.g., health literacy, history of physical or psychologic trauma, trust in the health system), which may influence clinical outcomes, were not available. Although older age has been identified as a potential effect modifier [21, 61], our sample could not assess this due to enrolling only adults under 65. We excluded individuals who reported a previous radiculopathy and those with an active or planned workers' compensation claim. Thus, we were unable to assess these potentially influential factors as predictors of response or effect modifiers.

Although loss to follow-up was generally low at the end of the 12-week intervention period (7%), another limitation of our study was the disproportionately high loss to followup in the PT arm (12%). This differential loss to follow-up may bias the results seen here, though the direction of this bias is unclear. Previous literature from this study has shown little difference between complete case and imputation-based analyses [24, 62]; therefore, we present only the findings for those with complete outcome data.

The strengths of our study include random allocation of adults to three credible nonpharmacologic cLBP treatments, and our extensive collection of baseline sociodemographic and health information from a diverse sample, which allowed us to explore a wide range of predictor variables. Lastly, we feel confident in our outcome definition, as it was prespecified [23], based on established clinically important differences for cLBP research [11, 12, 23], and aligned with recommendations from the NIH Task Force on Research Standards for Chronic Low Back Pain [10].

Current clinical practice guidelines suggest more than a dozen nonpharmacologic approaches as first-line treatment for cLBP [1]. Use of large data sets, such as claims data or practice-based research networks, may allow for additional opportunities to identify potential effect modifiers that can be used in shared decision-making for managing cLBP. Uniform collection of a minimal data set [10] in future cLBP clinical trials would allow for patient-level meta-analyses that may provide adequate statistical power to explore treatment effect modification across multiple guideline-concordant treatments. A lack of uniform information on baseline characteristics in clinical trials was cited as a key limitation in two recent individual patient-level data meta-analyses exploring effect modification of acupuncture and exercise interventions for low back pain [51, 63].

In conclusion, our findings revealed two important subgroups where referral to yoga or PT may be more likely to improve outcomes compared with an evidencebased self-care book. Adults using pain medications and those with few fear avoidance beliefs around physical activity were more likely to have a clinically meaningful response to yoga or PT compared with self-care. Although yoga and PT appear to be similarly effective, treatment expectations may influence clinical outcomes.

Authors' Contributions

Drs. Roseen, Gerlovin, and Saper had full access to all of the data in the study and take responsibility for the integrity of the data and the accuracy of the data analysis. Study concept and design: Roseen, Sherman, Delitto, Saper. Acquisition, analysis, or interpretation of data: Roseen, Gerlovin, Felson, Delitto, Sherman, Saper. Drafting of the manuscript: Roseen. Critical revision of the manuscript for important intellectual content: Roseen, Gerlovin, Felson, Delitto, Sherman, Saper. Statistical analysis: Roseen, Gerlovin. Obtained funding: N/A. Administrative, technical, or material support: Roseen, Saper. Study supervision: Roseen, Saper.

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Appendices



Appendix Figure 1. Clinical decision-making tree.

Appendix Table 1. Baseline characteristics* of analytic sample

	Total	Yoga	РТ	Self-care
Baseline Characteristics	(N = 299)	(N = 125)	(N = 113)	(N = 61)
Sociodemographic characteristics				
Age, mean \pm SD, y	46.1 ± 10.7	46.7 ± 10.3	46.4 ± 11.5	44.3 ± 9.97
18-44	117 (39.1)	50 (40)	41 (36.3)	26 (42.6)
45-65	182 (60.9)	75 (60)	72 (63.7)	35 (57.4)
Female	196 (65.6)	72 (57.6)	82 (72.6)	42 (68.9)
Race, nonwhite	55 (18.4)	29 (23.2)	16 (14.2)	16 (16.4)
Ethnicity, Hispanic	40/295 (13.6)	20/124 (16.1)	14/111 (12.6)	6/60 (10)
Education, >high school	180/296 (60.8)	74/123 (60.2)	63/112 (56.3)	43/61 (70.5)
Currently employed	135 (45.2)	59 (47.2)	47 (41.6)	29 (47.5)
Low annual income, ≤\$30,000	175/274 (63.9)	75/118 (63.6)	62/100 (62.0)	38/56 (67.9)
General health characteristics				
BMI, mean \pm SD, kg/m ²	31.8 ± 7.4	30.8 ± 6.7	32.7 ± 7.6	32.4 ± 8.1
Obese, BMI $> 30 \text{ kg/m}^2$	163 (54.5)	67 (53.6)	63 (55.8)	33 (54.1)
Comorbid chronic conditions, ≥ 2	141 (47.2)	53 (42.4)	55 (48.7)	33 (54.1)
Smoked cigarettes, any, in last week	91 (30.4)	36 (28.8)	37 (32.7)	18 (29.5)
Back pain measures				
Back pain intensity (0–10), mean \pm SD	7.2 ± 1.4	7.1 ± 1.4	7.3 ± 1.4	7.1 ± 1.4
Moderate, 4–6	169 (56.5)	71 (56.8)	60 (53.1)	68 (62.3)
Severe, 7–10	130 (43.5)	54 (43.2)	53 (46.9)	23 (37.7)
Back pain frequency, last 90 d, median	75	80	75	75
Limiting back pain, past 30 d, median	15	15	15	15
Pain medication use, any	210 (70.2)	85 (68.0)	80 (70.8)	45 (73.8)
Psychological measures				
Depression symptoms, PHQ-8, score ≥ 10	102/294 (34.7)	41/123 (33.3)	41/112 (36.6)	20/59 (33.9)
Anxiety symptoms, GAD-7, score ≥ 10	87/295 (29.5)	36/125 (28.8)	33/110 (30.0)	18/60 (30.0)
Perceived stress scale, PSS-10 \geq 17	157/295 (53.2)	63/123 (51.2)	62/112 (55.4)	32/60 (53.3)
Fear Avoidance Beliefs Questionnaire				
Physical activity score >13	155/296 (52.4)	69/124 (55.7)	57/112 (50.9)	29/60 (48.3)
Work score >29	42/288 (14.6)	14/124 (11.3)	19/104 (18.3)	9/60 (15.0)
Sleep quality, poor sleep, PSQI >5	276 (92.3)	115 (92.0)	106 (93.8)	55 (90.2)
Catastrophizing, CSQ \geq 14	155/298 (52.0)	60/125 (48.0)	67/112 (59.8)	28/61 (45.9)
Pain self-efficacy, PSEQ <40	166/298 (55.7)	70/125 (56)	64/112 (57.1)	32/61 (52.5)
Expectations				
Yoga would be helpful	121 (40.5)	52 (41.6)	44 (38.9)	25 (41.0)
PT would be helpful	141 (47.2)	59 (47.2)	55 (48.7)	27 (44.3)
Self-care would be helpful	73/296 (24.7)	27/124 (21.8)	34/111 (30.6)	12/61 (19.7)
Concordance, received a "helpful" treatment	119 (39.8)	52 (41.6)	55 (48.7)	12 (19.7)

BMI = body mass index; CSQ = Coping Strategies Questionnaire; FABQ = Fear Avoidance Beliefs Questionnaire; GAD-7 = Generalized Anxiety Disorder 7-item; PHQ-8 = Patient Health Questionnaire; PSEQ = Pain Self-Efficacy Questionnaire; PSQI = Pittsburgh Sleep Quality Index; PSS-10 = 10-item Perceived Stress Scale; PT = physical therapy.

*Presented as No. (%, out of total No.), unless otherwise noted.

Appendix Table 2. Con	nparison between tl	hose with and without	complete data on	outcome at 12 weeks
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Characteristics	Participants with Complete Data (N=299)	Participants with Missing Data*(N=21)		
Age, mean \pm SD, y	46.1 ± 10.7	44 ± 11.2		
Female,** No. (%)	196 (65.6)	8 (38.1)		
Race, nonwhite, No. (%)	244 (81.6)	18 (85.7)		
Ethnicity, Hispanic, No. (%)	40/295 (13.6)	3/21 (14.3)		
Education, >high school, No. (%)	180/296 (60.8)	11/21 (52.4)		
Currently employed, No. (%)	135 (45.2)	9 (42.9)		
Low annual Income, \leq \$30,000, No. (%)	175 (63.9)	13 (65)		
Back pain intensity (0–10), mean \pm SD	7.1 ± 1.4	7 ± 1.8		
Roland Morris, mean ± SD	14.8 ± 5.3	15.5 ± 6.3		

*Individuals missing data on Roland Morris Disability Questionnaire at 12 weeks were excluded from analyses.

**Statistically significant P = 0.01.

Appendix Table 3. Baseline characteristics of 299 participants, stratified by treatment and responder status

Baseline Characteristics	Total (N = 299)	Responders, No. (%)	P Value	Yoga (N = 125)	Responders, No. (%)	PT (N=113)	Responders, No. (%)	Self-care $(N = 61)$	Responders, No. (%)
Age			0.0009						
18–44 y	117	59 (50)		50	29 (58.0)	41	20 (49)	26	10 (39)
45–65 y	182	57 (31)		75	31 (41.3)	72	22 (31)	35	4 (11)
Sex			0.46						
Male	103	37 (36)		53	22 (42)	31	11 (35)	19	4 (21)
Female	196	79 (40)		72	38 (53)	82	31 (38)	42	10 (24)
Race			0.41						
Nonwhite	244	92 (38)		96	45 (47)	97	35 (36)	51	12 (24)
White	55	24 (44)		29	15 (52)	16	7 (44)	10	2 (20.0)
Ethnicity			0.89						
Non-Hispanic	255	99 (39)		104	48 (46)	97	38 (39)	54	13 (24)
Hispanic	40	16 (40)		20	11 (55)	14	4 (29)	6	1 (16.7)
Education	44.6	22 (20)	0.003	40	40 (20)	10	12 (27)	10	0 (0)
\leq High school	116	32 (28)		49	19 (39)	49	13 (27)	18	0 (0)
>High school	180	81 (45)	0.0001	/4	39 (53)	63	28 (44)	43	14 (33)
Employed	1.64	46 (20)	< 0.0001		22 (22)		10 (20)	22	5 (1 ()
No	164	46 (28)		66	22 (33)	66	19 (29)	32	5 (16)
Yes	135	/0 (52)	0.0002	59	38 (64)	4/	23 (49)	29	9 (31)
Annual income	175	54 (21)	0.0002	7.5	20 (27)	(2)	20 (22)	20	c(1c)
≤\$30,000 ≥ 20,000	1/3	54 (31)		/3	28(37)	62	20 (32)	38	6 (16) 7 (20)
>30,000	99	53 (54)	0.1.4	43	27 (63)	38	19 (50)	18	7 (39)
BMI categories	120	50 (42)	0.14	50	21 (52)	51	22/42)	20	((21)
Nonobese	136	59 (43)		38	31 (33)	51	22(43)	28	6 (21)
Obese	165	57 (33)	-0.0001	67	29 (43)	62	20 (32)	33	8 (24)
	150	79 (50)	<0.0001	72	12 (59)	50	26 (45)	20	11 (20)
0-1 >2	130	79 (30)		52	42 (38)	55	26 (43)	20	$\frac{11}{2}(32)$
<u>Z</u>	141	37 (28)	0.008	33	18 (34)	33	16 (29)	33	3 (9)
No	187	81 (13)	0.008	80	44 (55)	70	27 (39)	37	10(27)
Vec	107	29 (28)		41	13 (32)	41	$\frac{27}{32}$	23	3(13)
Coronary heart disease	105	27 (28)	0.49*	71	15 (52)	71	15 (52)	25	5 (15)
No	284	113 (40)	0.47	117	59 (50)	107	40 (37)	60	14 (23)
Yes	8	2 (25)		5	1(20)	2	1(50)	1	0(0)
Osteoarthritis	0	2 (23)	0.05	5	1 (20)	-	1 (50)	1	0 (0)
No	231	98 (42)	0.00	100	50 (50)	84	35 (42)	47	13 (28)
Yes	42	11 (26)		12	5 (42)	20	5 (2.5)	10	1(10)
Neck pain		()	0.04		- (/		- ()		- ()
No	197	83 (42)		78	41 (53)	80	31 (39)	39	11 (28)
Yes	89	26 (29)		40	16 (40)	27	7 (26)	22	3 (14)
Fibromyalgia		· · · ·	0.02		()		· · ·		()
Yes	17	2 (12)		8	1 (13)	7	1 (14)	2	0 (0)
No	270	112 (41)		111	58 (52)	103	41 (40)	56	13 (23)
Depression			0.0002						
No	225	102 (45)		102	53 (52)	77	36 (47)	46	13 (28)
Yes	63	12 (19)		20	7 (35)	31	5 (16)	12	0 (0)
Psychiatric disorder			0.14						
No	244	99 (41)		107	52 (49)	89	36 (40)	48	11 (23)
Yes	48	14 (29)		16	7 (44)	21	6 (29)	11	1 (9)
Hepatitis			0.32						
No	283	112 (40)		120	58 (48)	105	40 (38)	58	14 (24)
Yes	15	4 (27)		5	2 (40)	7	2 (29)	3	0 (0)
Diabetes			0.32						
No	237	96 (4)		99	49 (49)	88	34 (39)	50	13 (26)
Yes	57	19 (33)		25	11 (44)	24	8 (33)	8	0 (0)
COPD			0.13						
No	224	93 (42)		94	46 (49)	85	35 (41)	45	12 (27)
Yes	73	23 (32)		30	14 (47)	27	7 (26)	16	2 (13)
Cancer			0.33*						
No	288	114 (40)		122	59 (48)	107	41 (38)	59	14 (24)
Yes	10	2 (20)		2	1 (50)	6	1 (17)	2	0 (0)

(continued)

Appendix Table 3. continued

Baseline Characteristics	Total (N = 299)	Responders, No. (%)	P Value	Yoga (N = 125)	Responders, No. (%)	PT (N = 113)	Responders, No. (%)	Self-care (N=61)	Responders, No. (%)
Exercised last week			0.16						
No	102	33 (32)		35	11 (31)	44	17 (39)	23	5 (22)
Yes	118	49 (42)		55	28 (51)	43	15 (35)	20	6 (30)
Current smoker			< 0.0001						
No	208	97 (47)		89	50 (56)	76	33 (43)	43	14 (33)
Yes	91	19 (21)		36	10 (28)	37	9 (24)	18	0 (0)
Function (RMDQ)			0.04						
Mild (0-8)	45	21 (47)		28	17 (61)	7	2 (29)	9	2 (22)
Moderate (9–16)	129	57 (44)		49	24 (49)	52	23 (44)	29	10 (34)
Severe (17–24)	125	38 (30)		48	19 (40)	54	17 (31)	23	2 (9)
Pain			0.08						
Moderate (4–6)	169	73 (43)		71	34 (48)	60	27 (45)	38	12 (32)
Severe (7–10)	130	43 (33)		54	26 (48)	53	15 (28)	23	2 (9)
LBP every day, 3 mo			0.09						
No	196	83 (42)		78	44 (56)	75	27 (36)	43	12 (28)
Yes	102	33 (32)		46	16 (35)	38	15 (39)	18	2 (11)
Activity lim. daily, 1 mo			0.02						
No	244	102 (42)		100	53 (53)	91	36 (40)	53	13 (25)
Yes	55	14 (25)		25	7 (28)	22	6 (27)	8	1 (13)
Using pain medication			0.0005†						
None	89	48 (54)		40	24 (60)	33	15 (45)	16	9 (56)
Any	210	68 (32)		85	36 (42)	80	27 (34)	45	5(11)
NSAIDs			0.35†						
No	142	59 (42)		63	27 (43)	53	22 (42)	26	10 (38)
Yes	157	57 (36)		62	33 (53)	60	20 (33)	35	4 (11)
Acetaminophen		· · · ·	0.002†		()		· · ·		· · · ·
No	197	89 (45)		83	47 (57)	78	30 (38)	36	12 (33)
Yes	102	27 (26)		42	13 (31)	35	12 (34)	25	2 (8)
Muscle relaxant		· · · ·	0.04		()		· · ·		()
No	274	111 (41)		119	59 (50)	101	39 (39)	54	13 (24)
Yes	25	5 (20)		6	1 (17)	12	3 (25)	7	1 (14)
Opioids		()	0.005		· · ·		· · /		
No	251	106 (42)		104	55 (53)	96	37 (39)	51	14 (27)
Yes	48	10 (21)		21	5 (24)	17	5 (29)	10	0 (0)
FABO physical activity			$0.60 \pm$						
Low-risk (<13)	141	56 (40)	·	55	29 (53)	55	23 (42)	31	4 (13)
High-risk (>13)	155	57 (37)		69	30 (43)	57	18 (32)	29	9 (31)
FABQ Work		· · · ·	0.001		()		· · ·		· · · ·
Low-risk (<29)	246	105 (43)		110	58 (53)	85	34 (40)	51	13 (25)
High-risk (>29)	42	7 (17)		14	2 (14)	19	4 (21)	9	1 (11)
Depressive symptoms, PHQ-	8		< 0.0001		()		· · /		()
Low (<10)	192	93 (48)		82	47 (57)	71	33 (46)	39	13 (33)
High (>10)	102	23 (23)		41	13 (32)	41	9 (22)	20	1 (5)
Anxiety symptoms, GAD-7		· · · ·	0.29		(<i>'</i>		· · /		()
Low (<10)	208	83 (40)		89	45 (51)	77	27 (35)	42	11 (26)
High (>10)	87	29 (33)		36	15 (42)	33	12 (36)	18	2 (11)
Sleep quality, PSOI			0.002				(0 0)		- ()
Normal (<5)	23	14 (61)		10	6 (60)	7	6 (86)	6	2 (33)
Poor (>5)	276	102 (37)		115	54 (47)	106	36 (34)	55	12 (22)
PSS-10			0.004				(-)		()
Normal (< 17)	138	64 (46)		60	33 (55)	.50	23 (46)	28	8 (29)
Poor (≥ 17)	157	50 (32)		63	26 (41)	62	18 (29)	32	6 (19)
Catastrophizing, CSO	107	00 (02)	0.23	00	20 (11)	02	10 (=>)		0 (1))
Normal (<14)	143	61 (43)		65	33 (51)	45	19 (42)	33	9 (27)
High (>14)	155	55 (35)		60	27 (45)	67	23 (34)	28	5(18)
Self-efficacy. PSEO	100	23 (33)	0.001	00	=/ (13)	07	_3 (31)	20	5 (10)
Low/moderate (<40)	166	51 (31)	0.001	70	27 (39)	64	17 (27)	32	7 (22)
High (>40)	132	65 (49)		55	33 (60)	48	2.5 (52)	29	7 (24)
Expectations, voga	102		0.14	55	00 (00)	10	_3 (32)		/ (21)
Low/moderate. <9	178	63 (35)	V.T I	73	28 (38)	69	25 (36)	36	10 (28)
High. 9–10	12.1	53 (44)		52	32 (62)	44	17 (39)	2.5	4 (16)
		~~ (• • • /		~-	(/		(0) /	_0	. (10)

(continued)

Baseline	Total	Responders,	P Value	Yoga (N - 125)	Responders,	PT (N - 112)	Responders,	Self-care $(N - 61)$	Responders,
Characteristics	(1N - 299)	100. (70)	I value	(1N - 123)	100. (70)	(1N - 113)	110. (70)	(1N - 01)	100. (70)
Expectations, PT			0.58						
Low/moderate, <9	158	59 (37)		66	32 (48)	58	18 (31)	34	9 (26)
High, 9–10	141	57 (40)		59	28 (47)	55	24 (44)	27	5 (19)
Expectations, self-care			0.08						
Low/moderate, <9	223	93 (42)		97	50 (52)	77	29 (38)	49	14 (29)
High, 9–10	73	22 (30)		27	10 (37)	34	12 (35)	12	0 (0)
Expectations, concordance			0.02						
No	180	60 (33)		73	28 (38)	58	18 (31)	49	14 (29)
Yes	119	56 (47)		52	32 (62)	55	24 (44)	12	0 (0)

Concordance: high expectations of treatment received.

BMI = body mass index; COPD = chronic obstructive pulmonary disease; CSQ = Coping Strategies Questionnaire; FABQ = Fear Avoidance Beliefs Questionnaire; GAD-7 = Generalized Anxiety Disorder 7-item; PHQ-8 = Patient Health Questionnaire; PSEQ = Pain Self-Efficacy Questionnaire; PSQI = Pittsburgh Sleep Quality Index; PSS-10 = 10-item Perceived Stress Scale; PT = physical therapy; RMDQ = Roland Morris Disability Questionnaire.

 $^{*}P$ values calculated using Fisher's adjustment due to ${>}20\%$ of expected cell counts ${<}5.$

 $^{\dagger}\mbox{Indicates}$ effect modification by treatment group.