

# The Causal Role of Fatigue in the Stress-Perceived Health Relationship: A MetroNet Study

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**Background:** We conducted a cross-sectional survey of 4 primary care MetroNet centers in metropolitan Detroit. Our objective was to describe the causal role of fatigue in the relationship among stress, stress resiliency, and perceived health in primary care. Fatigue is a public health problem that has been linked to stress and poor health. The causal role of fatigue between stress and perceived health is unknown.

**Methods:** Four hundred surveys were distributed to adult patients in 4 primary care centers in metropolitan Detroit between 2006 and 2007. Internal consistency reliabilities and principal factor analyses were calculated for the key psychological scales. Perceived health is the primary outcome. Path models were used to study the relationship among stress, fatigue, and perceived health. We also modeled the impact of select stress resiliency factors including sleep, recovery, and social support.

**Results:** Of the 400 distributed surveys, 315 (78.7%) had a response rate of 70% or more and were included in the analysis. Respondents were predominantly middle aged (median age, 43 years); female (58.7%); and African American (52.0%). The majority worked full time (56.5%); did not have a college degree (77.7%); and were not married (55.2%). Fatigue was reported by 59% of respondents, 42.7% of which was unexplained. The path model supported the causal role of fatigue between stress and perceived health. The positive effects of sleep, recovery, and social support on fatigue, stress, and perceived health were validated.

**Conclusion:** Fatigue was common in this metropolitan primary care environment and completely mediated the relationship between stress and poor perceived health. Therefore, stress, when significant enough to cause fatigue, may lead to poor health. (J Am Board Fam Med 2010;23:212–219.)

**Keywords:** Fatigue, Stress, Perceived Health

Fatigue is a common and costly public health problem with well-recognized health, safety, and performance implications.<sup>1–3</sup> It affects up to 40% of the population and is a frequent complaint by primary care patients.<sup>4,5</sup> The subjective nature of fatigue coupled with the frequent lack of medical explanations renders it a true clinical challenge for primary care providers.<sup>4</sup> As with other medically unexplained symptoms,<sup>6</sup> stress is frequently impli-

cated in medically unexplained fatigue and in diseases hallmarked by fatigue, such as fibromyalgia and chronic fatigue syndrome.<sup>7–9</sup> In addition, essential components of stress resiliency, such as social support, sleep, and recovery, are inversely related to fatigue.<sup>10</sup> The detrimental effect of stress on health has been established, as have the stress implications of poor health.<sup>11–15</sup> Therefore, fatigue is an integral part of the intertwined stress-health relationship. However, the causal role of fatigue in this relationship is unknown.

We tested the causal role of fatigue in the relationship between stress and perceived health in a cross-sectional sample of attendants to 4 primary care centers in metropolitan Detroit. This sample allowed us to model stress and fatigue in different primary care settings where both problems are common. We focused not only on stressors but, importantly, also on possible stress resiliency factors, including sleep, recovery, and social support. This offered us the op-

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portunity to model a more ecologically relevant data set.

Perceived health was our primary outcome because it is a validated predictor of future morbidity and mortality<sup>16–19</sup> and a useful and feasible clinical tool.<sup>20,21</sup> This study will help advance the understanding of the complex relationship between stress, fatigue, and perceived health in primary care.

## Methods

### Procedure

After obtaining the appropriate institutional ethical approval from Wayne State University, 400 self-administered, anonymous questionnaires were distributed by a designated front desk person or a research assistant at 4 participating primary care centers between November 2006 and December 2007. These centers were all part of MetroNet, a practice-based research network in metropolitan Detroit.<sup>22</sup> MetroNet is a consortium of 24 primary care practices and was established in 2001 to advance practice-based research focused primarily on underserved patient populations. The 4 MetroNet practices were selected because of the racial and socioeconomic diversity they represented. In each of the 4 centers, 100 surveys were offered to 100 consecutive patients. Each respondent was informed of the purpose of the survey with an introductory information sheet. If a patient chose not to participate, his or her survey was turned in blank and not offered to another patient. Returned surveys were collected in designated boxes.

### Survey Instrument

The survey consisted of 63 questions about select sociodemographic characteristics, and visual analog scale (VAS)-type items (0–100) that measured fatigue, stress, perceived health, sleep, recovery, and general symptoms. The VAS items were anchored with opposing terms such as “never” and “daily” or “not at all” and “very much so.” The stress and fatigue items have been previously validated in various combinations.<sup>23–25</sup> Internal consistency was assessed with Cronbach  $\alpha$  coefficient (all  $\alpha$  coefficients were  $\geq 0.80$ ) and dimensionality was assessed with confirmatory factor analysis using the principal components method and varimax rotation (all loaded on one factor that explained 60% or more of the variance). A score pertinent to each of these

constructs was generated by averaging the scores on the corresponding VAS items. Higher fatigue, stress, and general symptoms scores indicated more fatigue, stress, and general symptoms, respectively. Higher perceived health, recovery, and sleep scores indicated better perceived health, recovery, and sleep, respectively.

Fatigue was measured with 6 VAS items that reflected physical and mental fatigue. Two items measured physical fatigue: level of energy and exhaustion; 4 items measured mental fatigue<sup>26</sup>: moodiness, restlessness, irritation, and difficulty concentrating. These items were adapted from the Brief Fatigue Syndrome Scale and the mental energy subscale of the Quality Work Competence questionnaire, which are validated and have been used in a number of studies, including prospective, controlled intervention studies targeting mental energy as well as fatigue.<sup>24,25</sup> In agreement with previously suggested cutoff levels,<sup>24</sup> a fatigue score of  $\geq 40$  on a 100% scale indicated fatigue and a score of  $\geq 75$  indicated severe fatigue.

Perceived stress was measured with 4 VAS items that assessed the extent of present and past pressure and stress. Consistent with the suggested cutoff points for the fatigue scale, a score of  $\geq 40$  was indicative of stress and a score of  $\geq 75$  was indicative of severe stress.

Recovery was measured with 4 VAS items that assessed the ability to rest, relax, recharge, and mobilize extra strength when needed. Sleep quality was measured with 5 VAS items that assessed perceived sleep quality presently and during the past year, restorative sleep, and difficulty initiating or sustaining sleep. Social support was assessed with a single VAS item concerned with the perception of social support. Social support, sleep, and recovery were the selected stress resiliency factors.

Perceived health was measured with 2 VAS items concerned with general health perceptions presently and during the past year.<sup>19–21</sup> General symptoms were assessed with 8 VAS items measuring the frequency of headache, anxiety, dizziness, pain, altered appetite, gastrointestinal symptoms, and cardiac symptoms. Health status was determined by the number of diagnosed diseases (diabetes, hypertension, cancer, prolonged pain, asthma, allergies, and cardiovascular and mental disorders) and was categorized as none, 1 to 2, and  $>2$  current diseases.

For the purposes of the analysis, race was categorized as African American, white, and other. In the path model only whites and African Americans were included because of the small number and heterogeneity of the “other” race group. Exercise was categorized as “mostly idle,” “some regular exercise,” and “regular moderate exercise 2 to 3 times a week.” Body size was based on self-reported height and weight, which was converted to body mass index (BMI) and defined as normal (BMI, 18.5–24.9 kg/m<sup>2</sup>); overweight (BMI, 25.0–29.9 kg/m<sup>2</sup>); obese (BMI, 30.0–39.9 kg/m<sup>2</sup>); and morbidly obese (BMI, ≥40 kg/m<sup>2</sup>).

### Statistical Analysis

Multiple imputation methods using SPSS Multiple Imputation (version 17.0, SPSS, Inc., Chicago, IL) were used to manage the nonresponse items. Summary statistics for the main scales (stress, fatigue, sleep, and recovery) were compared using complete case analysis and multiple imputation analysis and revealed very close values (data not presented). Manifest path analysis using LISREL version 8.72 (Scientific Software International, Inc., Lincolnwood, IL) was performed to test the hypothesized causal role of fatigue in the relationship between stress and perceived health.

Model fit was assessed using the comparative fit index (CFI), root mean square error of approximation (RMSEA), and 90% CI for RMSEA. CFI >0.95 and RMSEA <0.06 indicated a good-fitting model.<sup>27</sup> Paths that were not significant at  $P \leq .05$  were removed from the model. Modifications to the model were based on both substantive clinical and statistical grounds. All  $P$  values were based on 2-sided tests of statistical significance, which was set at 0.05.

## Results

### Study Population

Of the 400 distributed surveys, 65 were either unreturned or returned blank (16.2%) and 20 were missing >30% of the responses (5%) and were excluded from the analysis. Three hundred and fifteen patients answered >70% of the questions and were included in the analysis (n = 315; 78.7%). Of the 315, 219 (69.5%) had complete data and only 5.7% of the data were missing.

Table 1 summarizes the characteristics of the 315 respondents. Respondents tended to be mid-

**Table 1. Characteristics of Primary Care Patients in Metropolitan Detroit (n = 315)**

Characteristic	n (%)
Female	185 (58.7)
Age, years (mean, SD)	44.6 (14.9)
Race	
White	127 (40.3)
Black	164 (52.0)
Other	24 (7.6)
Education	
≤High school	104 (33.0)
Some college	133 (42.2)
≥College degree	70 (22.2)
Marital status	
Cohabiting	141 (44.7)
Single	106 (33.6)
Widowed/divorced	68 (21.5)
Employment status	
Unemployed	98 (31.1)
Part time	38 (12.0)
Full time	178 (56.5)
Body mass index	
Normal	75 (23.8)
Overweight	110 (34.9)
Obese	94 (29.8)
Morbidly obese	36 (11.4)
Exercise	
None	82 (26.0)
Some regular exercise	137 (43.4)
Regular moderate exercise	96 (30.4)
Current smokers	95 (30.1)
Diseases present	
0	148 (46.9)
1–2	135 (42.8)
>2	32 (10.1)
Fatigue cases*	186 (59.0)
Stress cases <sup>†</sup>	213 (67.6)

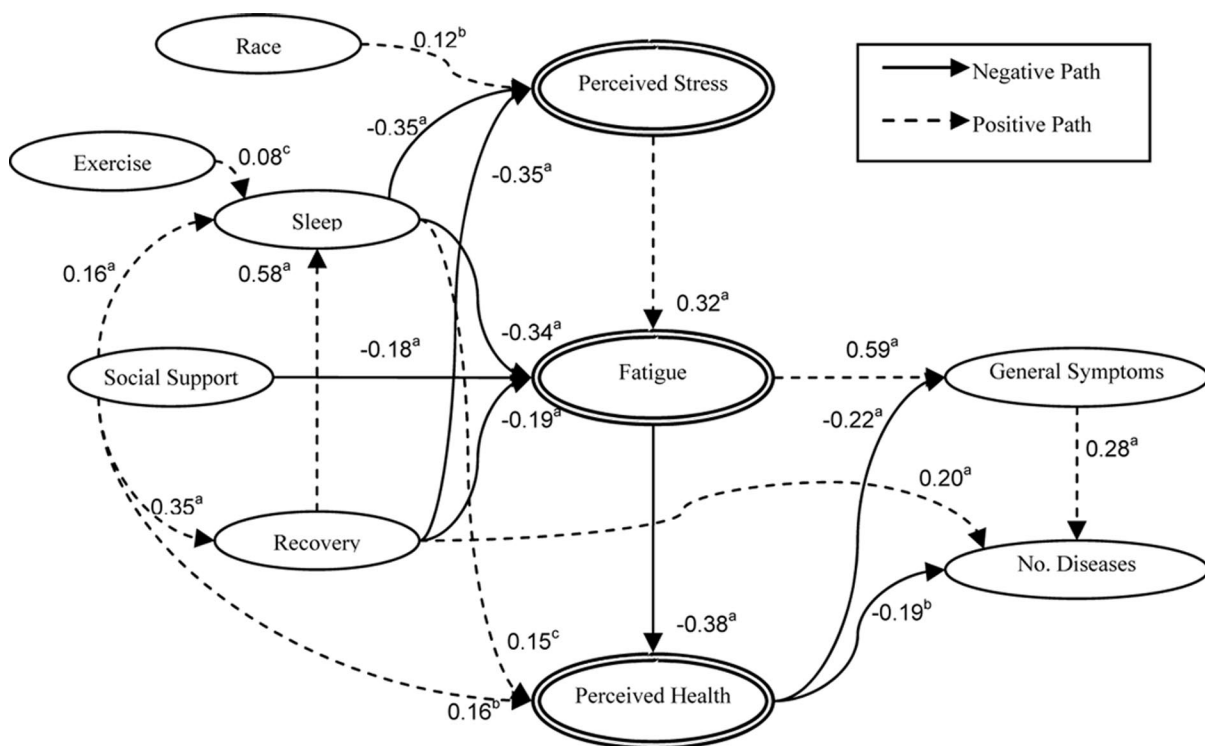
\*A fatigue case is defined by a fatigue score of ≥40.

<sup>†</sup>A stress case is defined by a perceived stress score of ≥40.

dle-aged (median age, 43 years; mean age, 44.6 years). More than half were women (n = 185; 58.7%) and the majority did not have college degree (n = 245; 77.7%). Close to half were either married or living with a significant other (n = 141; 44.7%). A little more than half identified themselves as African American (n = 164; 52.0%) and 40% identified as white (n = 127; 40.3%). The remaining patients were Native American (n = 4), Asian (n = 5), and other (n = 15).

The majority were working, either full time (n = 178; 56.5%) or part time (n = 38; 12.0%). Part-

**Figure 1.** A path model analysis showing the relationship between stress, fatigue, and perceived health. The standardized coefficients are reported on the corresponding arrows. <sup>a</sup>*P* < .001; <sup>b</sup>*P* < .01; <sup>c</sup>*P* < .05



time workers' views of their work were evenly split between positive (n = 16; 41.0%) and negative (n = 17; 43.5%) whereas full-time workers held predominantly positive views (n = 108; 60.6%).

The vast majority of respondents were overweight (n = 240; 76.1%), and 11% were morbidly obese (n = 36; 11.4%). Approximately a third were current smokers (n = 95; 30.1%). Thirty percent of respondents reported regular moderate exercise at least twice a week (n = 96; 30.4%).

Fifty-three percent reported one or more present diagnosed diseases (n = 167; 53.0%), with hypertension being the most common (n = 74; 23.4%), followed by prolonged pain (n = 55; 17.4%), allergic disorders (n = 49; 15.5%), and mental disorders (n = 43; 13.6%). Twenty-seven percent had 2 or more medical problems (n = 86; 27.3%).

Stress and fatigue were common. Fatigue was present in 59.0% of respondents (n = 186) and severe fatigue was present in 17% (n = 55, 17.4%). Forty-two percent of those fatigued had no medical or psychiatric illness (n = 79; 42.7%). Stress was present in 67.6% of respondents. Severe stress was present in 30% of respondents (n = 94; 29.8%).

**Path Analysis**

Figure 1 presents the final path model. The arrows indicate the direct effects between variables and the regression coefficients indicate the magnitude and the strength of these effects. The model fit the data well ( $\chi^2 = 37.63$  [23, n = 291]; *P* = .028; RMSEA, 0.045; 90% CI, 0.01–0.07; CFI, 0.99).

**Direct Effects**

Perceived health was predicted by better sleep (*P* = .03), less fatigue (*P* ≤ .001), and better social support (*P* = .003). Sleep, recovery, and race had direct effects on perceived stress. White respondents had higher levels of perceived stress than their African-American counterparts (*p* = .007). Better sleep (*P* ≤ .001) and recovery (*P* ≤ .001) predicted lower perceived stress.

**Indirect Effects**

Perceived stress had significant indirect effects on perceived health (*P* ≤ .001), general symptoms (*P* ≤ .001), and number of diseases (*P* ≤ .001). Increased levels of perceived stress predicted lower perceived health and increased general symptoms and number of diseases.

Recovery had an indirect effect on the number of diseases ( $P \leq .001$ ) that was in the opposite direction of the direct effect ( $P = .001$ ). The total standardized effect of recovery on the number of diseases was insignificant ( $P = .61$ ).

## Discussion

This cross-sectional study of primary care patients supported the causal role of fatigue in the relationship between stress and perceived health. When stress is associated with fatigue, stress is associated with poor perceived health; when stress is not associated with fatigue, the association between stress and perceived health is not significant. The positive effects of sleep, recovery, and social support on stress, fatigue, and perceived health were validated.

We found a high prevalence of fatigue in our sample; 59.0% reported fatigue and 17.4% reported severe fatigue of at least 1 month's duration. Consistent with the relevant literature about fatigue, medical and psychiatric explanations were often lacking<sup>4,28</sup>; 42.7% of those fatigued had no medical or psychiatric explanation for their symptoms. These rates are higher than rates reported previously, which ranged between 2% and 38% depending on the case definition and the population studied.<sup>1,4,5,29-31</sup> Moreover, the prevalence of fatigue is sensitive to the chosen scale and threshold.<sup>32</sup> Nonetheless, unexplained prolonged fatigue is a common problem among this population. We used validated questions that reflected essential mental and physical fatigue experiences and had been proven sensitive in ascertaining improvement in fatigue in controlled prospective studies of anti-fatigue interventions targeting cognition, exercise, sleep, and nutrition.<sup>24,25</sup>

Stress has well-recognized deleterious health effects on diseases such as hypertension, as well as other cardiovascular and metabolic diseases.<sup>12,33-35</sup> Stress was reported by the majority of our sample (67.6%). The inclusion of select stress resiliency factors such as sleep, recovery, and social support is a unique aspect of this study that allowed a more comprehensive and relevant evaluation of the stress-health relation. Congruent with the relevant literature, sleep and recovery were inversely related to stress.<sup>36-38</sup>

In the present study, white respondents reported more stress than their African-American counterparts. This finding comes at odds with most of the

reported literature, which shows higher stress among African Americans compared with whites because of racism-related stress and challenging socioeconomic experiences.<sup>39-41</sup> However, the integration of several ethnic, racial, and religious minorities in metropolitan Detroit, where minority groups are no longer the exception but rather the true fabric of the society, may provide a deeper sense of belonging and therefore may partly explain this discrepancy. Moreover, even though we modeled important stress resiliency factors including sleep, recovery, and social support, other important factors, such as individual coping styles and participation in religious and voluntary organizations, were not measured.<sup>42,43</sup>

Fatigue played an important role in the relationship between stress and perceived health. It fully mediated this relationship, which could mean that stress may lead to poor health outcomes when it is significant enough to drain stress resiliency resources and manifest as fatigue. When stress is not related to fatigue there is no impact of stress on perceived health. Fatigue has been previously conceptualized as a warning sign, indicating harmful accumulation of stress.<sup>44</sup> Therefore, individual and contextual stressors should be assessed in patients presenting with fatigue where no clear explanation is ascertained. Appropriate individual- and aggregate-level stress management and resiliency enhancement approaches should be offered to avert the deterioration of perceived health.

Sleep predicted perceived health directly and indirectly through fatigue and stress. This agrees with the evidence linking sleep to stress,<sup>37,38,45</sup> fatigue,<sup>36,46</sup> and perceived health.<sup>47,48</sup> Therefore, improving sleep may lead to better perceived health independent of stress and fatigue, possibly through inflammatory<sup>49</sup> and neurobiological<sup>50</sup> pathways.

The benefits of social support are well established<sup>51-54</sup> and are highlighted in this study. In our sample, strong social support was predictive of positive health perceptions directly and indirectly through favorable effects on sleep, recovery, and fatigue. Interestingly, marital status and employment were not predictive of perceived health. Therefore, support garnered from social sources outside work and committed relationships has strong influence on health.

Perceived health could be easily ascertained by primary care providers by asking patients to rate their health: "How do you think your general

health is now?" and "How do you think your general health will be next year?" These simple questions can help identify patients who are a higher risk of future morbidity and mortality.<sup>17,18,55</sup> The inverse relationship between perceived health and the extent of general symptoms and diseases is plausible and supported by the literature.<sup>17,18,56,57</sup> However, the cross-sectional nature of this study limited our ability to test the causal associations. The other limitations of the study are the reliance on self-reports of diseases, height, weight, and employment and the fact that it is not population based. In addition, although depression was not independently ascertained, this should not compromise the results because even though depression and fatigue overlap, depression represents a more severe form and is probably present in patients with extreme fatigue scores. Furthermore, the fatigue scale was constructed similar to a validated scale that can differentiate between fatigue and depression.<sup>24</sup>

Nonetheless, this study has important strengths. It specifically targets a demographically and socioeconomically diverse primary care population. It is the first study, to our knowledge, that examines the causal role of fatigue between stress and health. The primary care focus of this study is an advantage because fatigue and stress are prevalent in primary care settings, where they often present in concert with other common and costly medically unexplained symptoms.<sup>6</sup> The scales used in this study are short and simple and could be easily incorporated into clinical encounters to screen patients for stress and fatigue or to monitor their response to treatment. Perceived health was an appropriate outcome measure for its simplicity, subjectivity, and proven validity and clinical utility.

## Conclusion

We were able in the present cross-sectional study of primary care patients to demonstrate the important causal role of fatigue between stress and perceived health and the favorable effects exhibited by sleep, recovery, and social support. Therefore, stress, when severe enough to deplete the recovery resources and cause fatigue, may lead to poor perceived health. Fatigue can signal unresolved stress and serve as a window of opportunity to intervene and avert the deterioration of health perceptions.

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