Characterizing effects of mild traumatic brain injury and posttraumatic stress disorder on balance impairments in blast-exposed servicemembers and Veterans using computerized posturography

Joanna R. Wares, PhD; Kathy W. Hoke, PhD; William Walker, MD; Laura Manning Franke, PhD; David X. Cifu, MD; William Carne, PhD; Cheryl Ford-Smith, PT, DPT, MS, NCS

1Department of Mathematics and Computer Science, University of Richmond, Richmond, VA; 2Department of Physical Medicine and Rehabilitation, Virginia Commonwealth University, Richmond, VA; 3Defense and Veterans Brain Injury Center, Department of Veterans Affairs, Richmond, VA; 4Department of Physical Therapy, Virginia Commonwealth University, Richmond, VA

Abstract—The high rate of blast exposures experienced by U.S. servicemembers (SMs) during the recent conflicts in Iraq and Afghanistan has resulted in frequent combat-related mild traumatic brain injuries (mTBIs). Dizziness and postural instability can persist after mTBI as a component of postconcussion syndrome, but also occur among the somatic complaints of posttraumatic stress disorder (PTSD). The goals of this study were to examine the use of computerized posturography testing (CPT) to objectively characterize chronic balance deficits after mTBI and to explore the utility of CPT in distinguishing between combat and blast-exposed participants with and without mTBI and PTSD. Data were analyzed from a subject pool of 166 combat-exposed SMs and Veterans who had a blast experience within the past 2 yr while deployed. Using nonparametric tests and measures of impairment, we found that balance was deficient in participants diagnosed with mTBI with posttraumatic amnesia (PTA) or PTSD versus those with neither and that deficits were amplified for participants with both diagnoses. In addition, unique deficiencies were found using CPT for individuals having isolated mTBI with PTA and isolated PTSD. Computerized balance assessment offers an objective technique to examine the physiologic effects and provide differentiation between participants with combat-associated mTBI and PTSD.

Key words: balance, balance impairment, blast exposure, computerized posturography, impairment, mild traumatic brain injury, posttraumatic amnesia, posttraumatic stress disorder, servicemembers, Veterans.

INTRODUCTION

In Operation Iraqi Freedom (OIF), Operation Enduring Freedom (OEF), and Operation New Dawn (OND), U.S. servicemembers (SMs) have been subjected to a high

Abbreviations: BESQ = Blast Experience Screening Questionnaire; COM = center of mass; COMP = composite score; CPT = computerized posturography testing; EQ = equation; DSM = Diagnostic and Statistical Manual of Mental Disorders; MCB = Marine Corps Base; MINI = Mini-International Neuropsychiatric Interview PTSD module; mTBI = mild traumatic brain injury; OEF = Operation Enduring Freedom; OIF = Operation Iraqi Freedom; OND = Operation New Dawn; PCL-C = PTSD Checklist, Civilian version; PCS = postconcussion syndrome; PTA = posttraumatic amnesia; PTSD = posttraumatic stress disorder; SM = servicemember; SOT = Sensory Organization Test; TBI = traumatic brain injury; VA = Department of Veterans Affairs; VAMC = VA Medical Center; VCU rCDI-B = Virginia Commonwealth University retrospective Concussion Diagnostic Interview-Blast version.

*Address all correspondence to Joanna R. Wares, PhD; Department of Mathematics and Computer Science, University of Richmond, 28 Westhampton Way, Richmond, VA 23173; 804-287-6564; fax: 804-287-6444.

Email: jware@richmond.edu

http://dx.doi.org/10.1682/JRRD.2014.08.0197
rate of blast exposures, with explosive munitions accounting for 78 percent of wounded in action cases, the highest proportion for any large-scale conflict [1]. Traumatic brain injury (TBI) is one of the consequences of these blast exposures and is considered the “signature wound” of the OIF/OEF/OND conflicts. Among deployed SMs and Veterans, 19 percent are estimated to have sustained a TBI [2]. Mild TBI (mTBI), or concussion, is by far the most common category of TBI during these deployments, accounting for over 80 percent of cases [3], and up to 20,000 additional mTBIs occur in garrison annually [4]. Although indexed as mild based on initial severity, nearly 20 percent of those sustaining mTBI will develop postconcussion syndrome (PCS), a condition of persistent symptoms (≥3 mo) that may include physical, cognitive, and behavioral impairments [5–6].

Among the many potential consequences of TBI, the effects of imbalance are some of the more impactful on functional status, including capacity to return to work [7]. Balance, or postural stability, is defined as the ability to maintain the body’s center of gravity within the base of support with minimal postural sway [8]. To achieve balance, input from multiple sensory components—visual, proprioceptive, and vestibular—must be integrated and coordinated with the motor system via the cerebellum [9]. Subjective dizziness and postural instability (imbalance) are common acutely and chronically after moderate-to-severe TBI [10], as well as mTBI, where they can persist chronically as components of PCS [11]. In moderate-to-severe TBI, objective impairment of early balance function is ubiquitous, can be measured on routine physical examination, and is predictive of rehabilitation outcome [12–13]. Objective balance impairments persisting months to years after moderate-to-severe TBI have also been documented on computerized posturography testing (CPT), a method of quantifying balance through body weight shifts on a force plate under normal and altered sensory conditions [14]. The Sensory Organization Test (SOT) is a component of CPT that measures information about the integration of the sensory components of standing balance (visual, proprioceptive, and vestibular) and generates an outcome measure called the equilibrium score that assesses the overall coordination of these systems to maintain standing posture [9].

In sports-related mTBI, objective balance deficits have been shown acutely (1 wk) and subacutely (up to 90 d) compared with both baseline and controls using either the SOT [15–16], the Balance Error Scoring System [17], or the center of mass (COM) during gait [18]. Findings on the SOT, COM, and Dizziness Handicap Inventory, a subjective measure of dizziness, have been shown to be significantly interrelated in the assessment of balance after TBI [19]. In one investigation, participants with acute mTBI had higher magnitudes of sway when deprived of accurate visual cues, despite having no gross visual or neurologic impairments, leading researchers to comment on the subtle complexity and need for central integration of the multisensory contributions to balance [20–21]. Importantly, however, in these mTBI studies, the differences typically resolved within the first several weeks to months [22–24]. While there are an increasing number of small studies showing static or dynamic balance deficits following sports-related mTBI in the acute (1 wk) and subacute (1–12 wk) periods [18,20–23], there are no large-scale, well-controlled sports or trauma studies measuring objective balance deficits after mTBI beyond this time frame [20].

Veterans and SMs with blast exposure and suspected mTBI also commonly report persistent dizziness, vertigo, clumsiness, and imbalance symptoms [11,25–26]. However, data on objective postural stability in this population are sparse and generally lacking controls. Vanderploeg et al. demonstrated impaired tandem gait in Vietnam war Veterans with chronic dizziness after mTBI compared with controls [27]. However, this study used archival data and self-identified mTBI, so the validity of the findings is unclear. In a case series of OIF/OEF/OND SMs with blast-associated mTBI, Hoffer et al. reported an 84 percent incidence of acute dizziness symptoms and a substantial portion had abnormal CPT-SOT scores more than 30 d after injury [28]. Similarly, a range of studies without control comparisons have reported that between 46 and 74 percent of symptomatic Veterans with blast-associated mTBI of even longer duration (months to years postinjury) had abnormal SOT scores [29–31].

The frequency of comorbidities associated with combat mTBI [32], which may also be linked to balance deficits, further complicates the identification of a consistent pattern of balance deficits corresponding to blast-related mTBI. One of the most significant unstudied potential confounders for military populations is posttraumatic stress disorder (PTSD), which is found in up to one-third of OIF/OEF/OND combatants [33]. Chronic dizziness is a common symptom among individuals with combat-associated PTSD [34]. Although published CPT data are lacking for PTSD, it is well documented that postural instability is associated with anxiety disorders in general [35].

In summary, the objectivity of assessment provided by CPT may offer a means of both identifying and monitoring recovery of individuals with mTBI-associated...
balance deficits. But the few published studies that examine balance impairments in SMs and Veterans with mTBI have lacked appropriate controls with a history of combat deployment and blast exposure and have not examined confounding factors such as PTSD. This raises questions about the confounding role of other combat and blast-related conditions in the findings to date, as well as the utility of objective CPT findings to either support the mTBI diagnosis or monitor recovery from mTBI. In this investigation, we sought to characterize balance deficits after combat blast exposure (with and without TBI and/or PTSD) and to address the utility of using CPT to differentiate blast-exposed individuals with no diagnosed injury, mTBI, PTSD, or co-occurring mTBI and PTSD. We hypothesized that there would be a unique pattern of balance deficits defined by CPT for individuals with chronic mTBI when compared with controls or individuals with PTSD.

METHODS

Participants

Participants were recruited via letters and advertisements and from ambulatory healthcare clinics at the Hunter Holmes McGuire Department of Veterans Affairs (VA) Medical Center (VAMC) in Richmond, Virginia; Fort Lee Army Base in Prince George County, Virginia; Quantico Marine Corps Base (MCB) in Prince William County, Virginia; and Camp Lejeune MCB in Jacksonville, North Carolina. SMs and Veterans were eligible if they had a blast experience within the past 2 yr while deployed in OIF/OEF/OND. After approval from the institutional review boards of each institution/facility and approval from the Department of Defense’s Human Research Protection Office, eligible participants who consented to inclusion in the study were evaluated. Blast experience was defined as having any of the following symptoms or experiences during or shortly after exposure to blast or explosion: dazed, confused, saw stars, headache, dizziness, irritability, memory gap (not remembering injury or injury period), hearing loss, abdominal pain, shortness of breath, struck by debris, knocked over or down, knocked into or against something, helmet damaged, or medically evacuated. Severe and moderate TBI were the only exclusion criteria and were defined as more than 30 min in coma, brain bleeding or blood clot (abnormal brain computed tomography scan), or none of first 24 or more hours after event remembered (posttraumatic amnesia [PTA] > 24 h). Therefore, participants either had blast exposure without sustaining TBI or had sustained at least one blast-related mTBI.

As part of a larger, prospective longitudinal investigation, all participants underwent a comprehensive baseline assessment to collect demographic information; medical history including injuries and care received during their military service; specifics of blast exposure, injury, care, and sequelae; and current symptoms and level of functioning. For this initial exploratory study, the associations between data on injury diagnoses, presence of mTBI and PTSD, and balance testing were analyzed. Based on the availability and time constraints and burden of the subjects in this longitudinal investigation, the baseline assessment was accomplished in one of two ways: standardized interviews or checklists/questionnaires. We categorized 107 participants with mTBI and/or PTSD using the Virginia Commonwealth University retrospective Concussion Diagnostic Interview-blast version (VCU rCDI-B) and the Mini-International Neuropsychiatric Interview PTSD module (MINI; version 6.0), respectively. Interview data were not available for the remaining 62 participants, and so mTBI and PTSD status for these participants was determined using the Blast Experience Screening Questionnaire (BESQ) and PTSD Checklist, Civilian version (PCL-C), respectively.

The VCU rCDI-B is a combined unstructured and fully structured interview designed to affirm the presence of a blast-associated mTBI, either with or without PTSD. For those with multiple blast-related experiences, the self-identified “worst” potential concussive event was selected for interview. The interview data were independently reviewed by five experienced TBI physicians who individually rated each blast exposure in reference to the Department of Defense/VA common definition for mTBI [36]. A consensus diagnosis was obtained for each participant based on the physician majority rating. The BESQ is a modified version of the Walter Reed Army Medical Center Blast Injury Questionnaire, which characterizes blast effects [37]. Similar to the VCU rCDI-B interview, an algorithm using the combination of the alteration of consciousness items from the BESQ was used on the noninterviewed subset that gave the peak kappa (k = 0.59, 91% correctly classified) versus the physician consensus among the interviewed subset. Further, based on our clinical experience and supporting data from the athletic mTBI literature [36], we hypothesized that those having mTBI with PTA would be most likely to experience long-term impairment commonly associated with blast injury. Using interview and BESQ data, we divided the participants with mTBI into
those with PTA and those without PTA. The diagnosis of mTBI with PTA is referred to in the analyses as “blast mTBI,” and the group with mTBI without PTA was combined with those diagnosed to have not sustained mTBI and referred to as “no blast mTBI.” The MINI is a validated, short, structured diagnostic interview based on Diagnostic and Statistical Manual of Mental Disorders (DSM)-IV and International Classification of Diseases-9th Revision criteria that was developed by psychiatrists and clinicians jointly in the United States and Europe [38]. Each participant’s PTSD diagnosis was determined using “relaxed” DSM-IV criteria that ignored the A2 criterion and simulates DSM-V [30]. The PCL-C is a validated and widely used measure of self-reported PTSD symptom severity [39–40]. It should be noted that the items and Likert values on the civilian version are identical to the military version; the civilian version was chosen to avoid assuming that all individuals’ most stressful life event was during combat. For the PCL-C, ≥58 was used to define PTSD because this cut-point gave the peak kappa value (k = 0.54, 81% correct classification rate) in analysis of its receiver operating characteristics versus the MINI within the 107 interviewed participants [39].

Outcome Measures

All participants underwent complete balance testing regardless of underlying injury or diagnosis, history of dizziness or imbalance, or current difficulties. Postural stability and balance were measured with CPT on a dual-plate force platform, the NeuroCom Smart Balance Master (NeuroCom International Inc; Clackamas, Oregon). The specific CPT given was the SOT, which generates equilibrium scores that compare the largest anterior-posterior movements of the subject over the trial with a theoretical limit for six sensory condition tasks. The sensory conditions were as follows: (1) eyes open with a fixed surface and visual surroundings, (2) eyes closed with a fixed surface, (3) eyes open with a fixed surface and sway referenced visual surroundings, (4) eyes open with a sway referenced surface and fixed visual field, (5) eyes closed with a sway referenced surface, and (6) eyes open with a sway referenced surface and visual surroundings (Figure 1). Each subject performed 3 trials on the Balance Master for each of the 6 sensory conditions, resulting in 18 equilibrium scores, ranging from 0 (touching a support surface, shifting feet, or falling) to 100 (little or no sway). From these equilibrium scores, 7 outcome measures were derived; the average of the 3 trials for each of the 6 conditions (equation [EQ] 1–EQ6) and an overall composite score (COMP) calculated as a weighted average of the 18 individual equilibrium scores (conditions 1 and 2 are weighted 1/3 as much as conditions 3 through 6).

Additionally, for this study “impairment” was defined as scoring at or below the 20th percentile of performance in a population of participants of similar age and with no history of disequilibrium (normative data provided by the administration manual).

Statistical Analyses

All statistical analyses were conducted using SPSS Statistics version 21.0 (IBM Corporation; Armonk, New York). Data were assessed for normality using the Shapiro-Wilk test. Since data were generally not normally distributed (or even transformed-normal), nonparametric Mann-Whitney U
and Kruskal-Wallis tests were used for between-group comparisons. When significant differences were found between groups, post hoc pairwise comparisons were performed using Dunn’s procedure with a Bonferroni correction for multiple comparisons. Mann-Whitney U tests were also performed on split data, and chi-square tests were applied to examine associations between participant cohorts and impairment.

RESULTS

Demographic Data

Of the 169 combat-exposed research participants, two participants’ data were removed due to missing outcome measures (both unable to tolerate test), and one participant’s data were removed because the balance scores did not pass the validity test for sufficient test effort (equilibrium scores for the more difficult conditions 5 or 6 were higher than for conditions 1, 2, or 3, pairwise) [41]. Of the 166 remaining participants with complete data, 160 were male. The mean age of the participants was 27.5 yr, with a standard deviation of 7.8 yr. Twenty-seven participants were African-American, 127 were white, and the remaining 12 self-identified as “other.” The median time since the self-identified “worst” potential concussive event was 11.6 mo, with an interquartile range of 13.7.

Of the 166 participants, 33 had no blast mTBI, 47 had blast mTBI without PTA, and 86 had blast mTBI with PTA. Forty-six were diagnosed with PTSD. For the purposes of data analysis, four subgroups were created: no diagnosis of PTSD or blast mTBI with PTA (n = 65), diagnosis of blast mTBI with PTA but not PTSD (n = 55), diagnosis of PTSD but not blast mTBI with PTA (n = 25), and diagnosis of both blast mTBI with PTA and PTSD (n = 21) (Table 1).

The SOT findings for all 166 participants with complete data were analyzed to characterize impairments on the seven outcome measures and to contrast findings between the cohorts.

No Blast mTBI Versus Blast mTBI (with PTA)

To explore whether individuals with blast mTBI exhibit balance deficits (regardless of the presence of PTSD), Mann-Whitney U tests were used to compare data for participants without blast mTBI (n = 80) and those with blast mTBI (n = 86) for each of the seven outcome measures. Only EQ3 showed a significant between-group difference (p = 0.006; no blast mTBI median = 92.3, interquartile range = 4.67; blast mTBI median = 90.5, interquartile range = 8.00).

Next, a chi-square test for each of the seven measures was used to test for association between impairment and blast mTBI diagnosis. A statistically significant association (p < 0.05) was found between blast mTBI diagnosis and impairment for COMP, as well as for EQ3 and EQ5 (Table 2).

PTSD Versus No PTSD

Similar analyses were performed for PTSD (regardless of the presence of blast mTBI). Mann-Whitney U tests were used to compare data for participants not diagnosed with PTSD (n = 120) and participants diagnosed with PTSD (n = 46) for each of the seven outcome measures. The Mann-Whitney U tests showed significant differences for the COMP (p = 0.01), EQ2 (p = 0.048), EQ4 (p = 0.007), EQ5 (p = 0.03), and EQ6 (p = 0.02) outcomes. In addition, chi-square analyses showed a significant association between impairment and PTSD according to the same five measures (Table 3).

Table 1.
Descriptive data by diagnosis.

<table>
<thead>
<tr>
<th>Group</th>
<th>Diagnosis</th>
<th>Male (n)</th>
<th>Female (n)</th>
<th>Age (yr), mean ± SD (n)</th>
<th>African American (n)</th>
<th>White (n)</th>
<th>Other Ethnicity (n)</th>
</tr>
</thead>
<tbody>
<tr>
<td>0 (n = 55)</td>
<td>No PTSD and No Blast mTBI with PTA</td>
<td>50</td>
<td>5</td>
<td>26.3 ± 7.6 (n)</td>
<td>9</td>
<td>43</td>
<td>3</td>
</tr>
<tr>
<td>1 (n = 65)</td>
<td>Blast mTBI with PTA only</td>
<td>65</td>
<td>0</td>
<td>27.5 ± 6.6 (n)</td>
<td>8</td>
<td>52</td>
<td>5</td>
</tr>
<tr>
<td>2 (n = 25)</td>
<td>PTSD only</td>
<td>24</td>
<td>1</td>
<td>29.0 ± 10.7 (n)</td>
<td>2</td>
<td>19</td>
<td>4</td>
</tr>
<tr>
<td>3 (n = 21)</td>
<td>Blast mTBI with PTA and PTSD</td>
<td>21</td>
<td>0</td>
<td>29.0 ± 7.8 (n)</td>
<td>8</td>
<td>13</td>
<td>0</td>
</tr>
</tbody>
</table>

mTBI = mild traumatic brain injury, PTA = posttraumatic amnesia, PTSD = posttraumatic stress disorder, SD = standard deviation.
Controls, Isolated Blast mTBI, Isolated PTSD, and Comorbid Blast mTBI/PTSD

Since 21 participants had both blast mTBI and PTSD, the interaction between mTBI and PTSD was investigated. First, Kruskal-Wallis tests were performed to determine whether there were differences in any of the equilibrium scores between the four mutually exclusive sets: participants diagnosed with neither blast mTBI nor PTSD (group 0; \( n \)= 55), participants diagnosed only with blast mTBI (group 1; \( n \)= 65), participants diagnosed with only PTSD (group 2; \( n \)= 25), and participants diagnosed with both blast mTBI and PTSD (group 3; \( n \)= 21).

The Kruskal-Wallis tests showed significant differences between groups for COMP, EQ3, EQ4, and EQ6 (see Figures 2 and 3 for results for COMP and EQ3.) Post hoc analyses indicated significant differences between groups 0 and 1 on EQ3, between groups 0 and 2 on EQ4, and between groups 0 and 3 on all four (COMP, EQ3, EQ4, and EQ6). There were no between-group differences found for groups 1, 2, and 3 in post hoc analyses.

To further investigate the interaction of blast mTBI and PTSD, individuals with comorbid conditions (group 3) were excluded and separate Mann-Whitney \( U \) tests comparing the data from participants having neither diagnosis (group 0) with participants having either isolated blast mTBI (group 1) or isolated PTSD (group 2) were performed. The same tests were then performed excluding group 0 and comparing group 3 with either group 1 or group 2 to determine whether co-occurring diagnoses would mask or amplify findings from the isolated injury groups.

Mann-Whitney \( U \) tests showed significant differences for COMP, EQ1, EQ2, EQ3, EQ4, and EQ6 when comparing group 0 (having neither diagnosis) to group 2 (isolated PTSD). In addition, using a chi-square measure of association, impairment had a significant association between group 0 and group 2 according to COMP, EQ2, EQ4, EQ5, and EQ6 (Table 4).

<table>
<thead>
<tr>
<th>Measure</th>
<th>Cut-Point</th>
<th>No Blast mTBI, % Impaired</th>
<th>Yes Blast mTBI, % Impaired</th>
<th>Chi-Square</th>
<th>( p )-Value*</th>
</tr>
</thead>
<tbody>
<tr>
<td>Composite Score</td>
<td>75</td>
<td>18.75</td>
<td>34.88</td>
<td>5.460</td>
<td>0.02</td>
</tr>
<tr>
<td>EQ1</td>
<td>92</td>
<td>21.25</td>
<td>24.42</td>
<td>0.236</td>
<td>0.63</td>
</tr>
<tr>
<td>EQ2</td>
<td>88</td>
<td>23.75</td>
<td>31.40</td>
<td>1.209</td>
<td>0.27</td>
</tr>
<tr>
<td>EQ3</td>
<td>88</td>
<td>20.00</td>
<td>33.72</td>
<td>3.949</td>
<td>0.047</td>
</tr>
<tr>
<td>EQ4</td>
<td>76</td>
<td>22.50</td>
<td>33.72</td>
<td>2.571</td>
<td>0.11</td>
</tr>
<tr>
<td>EQ5</td>
<td>60</td>
<td>15.00</td>
<td>27.91</td>
<td>4.065</td>
<td>0.04</td>
</tr>
<tr>
<td>EQ6</td>
<td>57</td>
<td>12.50</td>
<td>18.60</td>
<td>1.169</td>
<td>0.28</td>
</tr>
</tbody>
</table>

Note: Bold indicates statistical significance.
*All cells have expected count greater than 5.

EQ = equation.

Table 2.
Association between blast mild traumatic brain injury (mTBI) with posttraumatic amnesia and impairment on Sensory Organization Test.
In sharp contrast, Mann-Whitney $U$ tests showed no significant differential effect for PTSD when comparing data for participants diagnosed with isolated blast mTBI (group 1) to those with comorbid mTBI and PTSD (group 3). Likewise, impairment was not significantly associated with PTSD when only considering those with blast mTBI according on any of the seven measures (Table 4).

When the groupings were reversed and those having neither diagnosis (group 0) were compared with isolated blast mTBI (group 1), chi-square was significant for impairment according to COMP, EQ2, EQ3, EQ4, and EQ5 (Table 5). Again in sharp contrast, impairment was not significantly associated with mTBI when only considering those with PTSD according to any of the seven measures.

Finally, to determine whether participants exhibited differences between trials, Kruskal-Wallis tests were performed for groups 0 through 3 across the individual trials. No clinically significant results were found.

**DISCUSSION**

Traditionally, mTBI and PCS have been diagnosed by interview and physical examination proximate to the time of injury (e.g., in the emergency department); by interview and physical examination days, weeks, or months postinjury; and/or by neuroimaging. The ability to support the diagnosis of mTBI and assess the status of persistent difficulties from mTBI with physiological measures would improve the objectivity and reliability of diagnosis, allow for monitoring of recovery, and facilitate the assessment of treatment efficacy. The utility of assessing and defining balance deficits acutely after mTBI with CPT is well documented. Given the frequency of balance-related complaints and clinical findings following mTBI, identifying patterns of postural instability using CPT may represent a means of accurately identifying and quantifying the severity of balance deficits that may need treatment. CPT has also been suggested as an objective assessment tool for identifying and tracking the late effects of mTBI. However, previously chronically persisting balance deficits after combat blast mTBI have only been demonstrated in uncontrolled research studies and case reports. While the specific conditions and disorders that may contribute to postural instability may be of interest to patients and clinicians, this investigation focused on the use of the standardized SOT components of the Smart...
Table 4.
Association between posttraumatic stress disorder (PTSD) and balance (Sensory Organization Test) impairment.

<table>
<thead>
<tr>
<th>Measure</th>
<th>No PTSD, mTBI (n = 80)</th>
<th>Yes PTSD, mTBI (n = 86)</th>
</tr>
</thead>
<tbody>
<tr>
<td></td>
<td>Composite Score</td>
<td></td>
</tr>
<tr>
<td>No PTSD</td>
<td>10.91</td>
<td>42.86</td>
</tr>
<tr>
<td>PTSD</td>
<td>36.00</td>
<td>0778</td>
</tr>
<tr>
<td>Chi-Square</td>
<td>7.103</td>
<td>0.008</td>
</tr>
<tr>
<td>P-Value*</td>
<td></td>
<td></td>
</tr>
<tr>
<td>EQ1</td>
<td>16.36</td>
<td>24.62</td>
</tr>
<tr>
<td>EQ2</td>
<td>14.55</td>
<td>23.81</td>
</tr>
<tr>
<td>EQ3</td>
<td>14.55</td>
<td>33.33</td>
</tr>
<tr>
<td>EQ4</td>
<td>14.55</td>
<td>41.05</td>
</tr>
<tr>
<td>EQ5</td>
<td>7.27</td>
<td>42.86</td>
</tr>
<tr>
<td>EQ6</td>
<td>7.27</td>
<td>28.57</td>
</tr>
</tbody>
</table>

Note: Bold indicates statistical significance.
*All cells have expected count greater than 5.
EQ = equation, mTBI = mild traumatic brain injury.

Table 5.
Association between blast mild traumatic brain injury (mTBI) with posttraumatic amnesia and balance (Sensory Organization Test) impairment.

<table>
<thead>
<tr>
<th>Measure</th>
<th>PTSD = 0 (n = 120)</th>
<th>PTSD = 1 (n = 46)</th>
</tr>
</thead>
<tbody>
<tr>
<td></td>
<td>Composite Score</td>
<td></td>
</tr>
<tr>
<td>No PTSD</td>
<td>10.91</td>
<td>36.00</td>
</tr>
<tr>
<td>PTSD</td>
<td>32.31</td>
<td>42.86</td>
</tr>
<tr>
<td>Chi-Square</td>
<td>7.823</td>
<td>0.005</td>
</tr>
<tr>
<td>P-Value*</td>
<td></td>
<td></td>
</tr>
<tr>
<td>EQ1</td>
<td>16.36</td>
<td>32.00</td>
</tr>
<tr>
<td>EQ2</td>
<td>14.55</td>
<td>44.00</td>
</tr>
<tr>
<td>EQ3</td>
<td>14.55</td>
<td>32.00</td>
</tr>
<tr>
<td>EQ4</td>
<td>14.55</td>
<td>32.00</td>
</tr>
<tr>
<td>EQ5</td>
<td>7.27</td>
<td>32.00</td>
</tr>
<tr>
<td>EQ6</td>
<td>7.27</td>
<td>24.00</td>
</tr>
</tbody>
</table>

Note: Bold indicates statistical significance.
*No more than 1 cell per condition has expected count less than 5.
EQ = equation, PTSD = posttraumatic stress disorder.

Balance Master to differentiate individuals with diagnoses of mTBI and/or PTSD.

This is the first controlled study to examine the use of CPT to objectively characterize chronic balance deficits after combat-related mTBI and to explore the utility of CPT in distinguishing between combat and blast-exposed Veterans and SMs with and without mTBI and PTSD. Key findings of this investigation include (1) the characterization of balance deficits using CPT for participants having combat blast-associated mTBI with PTA or PTSD, (2) the confirmation of the amplification of CPT abnormalities in the face of both mTBI with PTA and PTSD, and (3) the identification of unique abnormalities on CPT for individuals with isolated mTBI with PTA or PTSD. These findings have potential implications for diagnostics, classifying residual mTBI-related impairments, and establishing treatment needs for mTBI and PTSD related postural instability.

CONCLUSIONS

No Blast mTBI with PTA Versus Blast mTBI with PTA

Balance performance on CPT of participants having blast mTBI with PTA differed significantly from blast-exposed controls having no mTBI or having mTBI without PTA, even in the context of high rates of PTSD. Uniformly,
when surveying a cohort of combat-exposed participants with and without PTSD, the median of condition 3 (sway-referenced visual surround) equilibrium scores of participants having mTBI with PTA was significantly lower than the group of controls having either no mTBI or having mTBI without PTA. Analyses of the incidence of balance impairment also revealed differences between these two groups for both the COMP and condition 5 equilibrium scores.

Taken together, these results provide evidence that, on average, persons with a history of blast mTBI with PTA have reduced postural stability relative to those without it. They corroborate findings from sports concussion cohorts showing mTBI with PTA has a poorer prognosis than mTBI without PTA [42]. As one would expect given the high incidence of visual tracking deficits seen with mTBI [43], the CPT conditions targeting inaccurate visual feedback (condition 3) or deprived visual and proprioceptive feedback (condition 5) were particularly sensitive to mTBI with PTA. On the contrary, when normal visual inputs were available as in conditions 1 and 4, regardless of the presence of normal or altered proprioceptive input (a sense infrequently affected by mTBI), there were no group differences. While PTSD did have effects on postural instability as seen on CPT, the effects from mTBI were still noted over the entire cohort, suggesting these findings are specific to mTBI with PTA. These findings also support the work of Vanderploeg et al. that indicated long-lasting gait deficits after mTBI [27] and agree with the other recent reports regarding Veterans and SMs that posturography is among the most consistently affected measures of the vestibular and balance system after blast injury [44] and blast mTBI [29–31]. The present findings extend these previous investigations with evidence that PTSD-related difficulties may not completely explain postural deficits in a military blast-exposed population. Additionally, the fact that these participants were not specifically referred for balance impairments or dizziness provides an unbiased perspective on balance deficits with PTSD and strengthens the generalizability of the findings.

**PTSD versus No PTSD**

Balance performance on CPT of participants with blast-exposed PTSD differed significantly from blast-exposed controls without PTSD, even in the context of mTBI. Uniformly, when surveying a cohort of combat-exposed participants with and without blast mTBI with PTA, the median COMP and condition 2, 4, 5, and 6 (all of the eyes closed or moving platform conditions) equilibrium scores of participants with PTSD were lower than those without PTSD. Taken together, these results provide evidence that persons with a history of PTSD have reduced postural stability relative to those without PTSD. As one would expect, given the diffuse effect of PTSD on attention, concentration, and the integration of sensory inputs on overall functioning, postural abnormalities on CPT were seen on almost all elements of testing. Impairments during both eyes open and closed suggest a multilevel deficit involving integration of vestibular, somatosensory, and visual information (i.e., the entire balance system). These findings echo those of the Jacob et al. study of anxiety disorders and SOT [35], but in contrast to the specific deficits noted in that study for spatial anxiety in panic and agoraphobic disorders (condition 4 only), PTSD appears to have a global effect on postural stability that is not indicative of an overreliance on a particular information channel. This could reflect a general attentional bias toward the “imbalance” signal—a mismatch between the gravitational vertical and other sensory inputs and a form of danger signal. While mTBI did have effects on postural instability as seen on CPT, the effects from PTSD were noted over the entire cohort, suggesting a distinctly different profile from mTBI. As stated previously, the fact that these participants were not specifically referred for balance impairments or dizziness provides an unbiased perspective on balance deficits with PTSD and strengthens the generalizability of the findings.

**Neither Diagnosis, Isolated Blast mTBI with PTA, Isolated PTSD, and Comorbid mTBI/PTSD**

Postural instability impairments are seen in participants having isolated blast mTBI with PTA or PTSD and are identifiable when compared with combat-exposed controls with neither diagnosis. As demonstrated in Tables 2 and 4, while there appear to be unique patterns of CPT findings of abnormalities for both mTBI (condition 3) and PTSD (conditions 2, 4, and 6), there are also overlapping abnormalities (condition 5 and the overall COMP). Unfortunately, for comorbid mTBI/PTSD, there appear to be only nominal abnormalities on the full range of scores compared with both isolated mTBI and PTSD that prevent the simple differentiation of the two conditions using CPT. In particular, when the investigation of balance deficits
after blast mTBI with PTA was limited to the subpopulation of participants with PTSD, no differences could be seen, suggesting PTSD masks the mTBI effects. The same result was found in the subpopulation of patients having blast mTBI with PTA: no median differences in scores were found between participants that were diagnosed with PTSD and those that were not because mTBI masked the effects of PTSD. Thus, the CPT findings may be useful to explain some percentage of the variance contributing to the differential diagnosis of mTBI and PTSD, but a more multimodal assessment tool may be needed to fully differentiate their effects.

Importantly, in individuals with both blast mTBI with PTA and PTSD, there is an overall amplification of abnormalities seen on CPT. Thus, individuals diagnosed with comorbid mTBI and PTSD would be expected to experience worse symptoms (dizziness), clinical findings (postural instability), and functional deficits (falls, inability to run) than those diagnosed with either of the conditions separately. Awareness of the cumulative effects of mTBI and PTSD on balance deficits may have clinical implications (e.g., earlier or more intensive use of vestibular rehabilitation). While a standard course of therapy to manage persistent postural instability due to distant blast mTBI with PTA may have a positive effect in individuals with isolated mTBI, clinicians may consider providing either a greater intensity or greater duration of these services in the face of concurrent PTSD; however, further research is needed to confirm this approach or to identify whether a different type of service may be needed altogether. While the specifics of therapeutic adjustments have not been elucidated, the ability of CPT to objectively (and differentially) identify abnormalities will assist in both developing and assessing the efficacy of these needed treatments.

Effective coordination of movement and balance involves a complex interaction of the sensory, motor, programming, and musculoskeletal systems. Even minor impairments in integrating this information could ostensibly lead to significant disability [45]. Persistent balance deficits, even if mild, can complicate recovery from brain injury by contributing to emotional distress. Even mild dizziness and balance problems are more highly associated with psychiatric comorbidity than other disturbances of sensory function, such as hearing loss [46]. This relationship is believed to be due to the closely shared neural circuitry between spatial processing, balance control, and arousal [47] and may in part explain why severe TBI patients with balance problems have a poorer prognosis than those without [12]. Many SMs and Veterans with mTBI balance deficits also have PTSD, and it is known that the combination of psychiatric and physical morbidities is particularly disabling [48]. Thus, it is important to treat balance deficits when they occur, particularly if the individual is at risk for developing an emotional disorder. Therapies successful in alleviating balance problems may reduce long-term disability and also have downstream benefits for emotional outcomes.

This large, prospectively collected sample did not have selection bias based on complaints of imbalance or dizziness and represents the first sizable cohort of individuals with combat-associated, chronic mTBI with either a comparison sample of combat-exposed controls or assessment of the confounding effects of PTSD. Despite the strengths of the study, some limitations should be noted: (1) a single data set from one medical center (albeit recruited from several military treatment centers and a VAMC), (2) an almost exclusive male population, (3) no controls for other potential contributors of balance deficits (explained subsequently), and (4) no “gold standard” for confirming the late diagnosis of chronic mTBI. Future studies using this data set and others should also assess other specific characteristics seen with combat trauma, mTBI, and cohorts of SMs that may contribute to balance deficits such as substance use, other neuropsychiatric disorders (e.g., depression, anxiety), neurosensory symptoms (diplopia, tinnitus, hearing loss), somatic symptoms (insomnia, limb numbness, pain), and structural impairments (inner ear damage, perilymph fistula, peripheral nerve injury). The findings of this investigation should also be crossvalidated in additional cohorts, including ones with significant female participants.

ACKNOWLEDGMENTS

Author Contributions:
Acquisition of data: W. Walker, L. Manning Franke.
Critical revision of manuscript for important intellectual content: D. X. Cifu, J. R. Wares, K. W. Hoke, L. Manning Franke, W. Walker, W. Carne, C. Ford-Smith.
Statistical analysis: K. W. Hoke, J. R. Wares.

Financial Disclosures: The authors have declared that no competing interests exist.

Funding/Support: This material was based upon work supported by the U.S. Army Medical Research and Materiel Command (award no. W81XWH-13-2-0095). The U.S. Army Medical Research Acquisition Activity, Fort Detrick, Maryland, is the awarding and administering acquisition office. This material was also based upon work supported with resources and the use of facilities at the Hunter Holmes McGuire VAMC in Richmond, Virginia, and work supported in part by the Defense and Veterans Brain Injury Center, U.S. Army Medical Research and Materiel Command.

Additional Contributions: The authors would like to thank University of Richmond-funded undergraduates Hilary Briggs, Ruoping Shi, WeiZhi Wu, and Xiwen Zhou for their exemplary research assistance on this project.

Institutional Review: The institutional review boards of each institution/ facility and the Department of Defense’s Human Research Protection Office approved this study.

Participant Follow-Up: The authors do not plan to inform participants of the publication of this study.

Disclaimer: The views, opinions, and/or findings contained in this article are those of the authors and should not be construed as an official VA or Department of Defense position, policy, or decision unless so designated by other official documentation.

REFERENCES


39. Walker WC, McDonald SD, Franke LM. Diagnostic accuracy of Posttraumatic Stress Disorder Checklist in blast

Submitted for publication August 26, 2014. Accepted in revised form April 1, 2015.

This article and any supplementary material should be cited as follows:

ResearcherID: Laura Manning Franke, PhD: F-2670-2012