#### Introduction

Premenstrual Dysphoric Disorder (PMDD) is characterized by extreme mood symptoms that emerge cyclically in the late luteal phase of the menstrual cycle and subside during menses. About 3-8% of women meet the full DSM-5 criteria<sup>1,2</sup> for PMDD, which requires the presentation of five symptoms (at least one being an emotional symptom) following the cyclical pattern per cycle, while another 10-11% of women can be characterized as having a less severe menstrually-related mood disorder (MRMD), which requires just one emotional symptom to follow the cyclical pattern per cycle<sup>2</sup>. Women with signs of MRMD have less severe emotional symptoms than those with PMDD. However, both PMDD and MRMD can cause impairment and distress severe enough to warrant treatment.

Diagnosis of PMDD or MRMD must follow strict criteria established by the DSM-5 over a period of at least two months of prospective daily symptom ratings. Since retrospective reports of these symptoms is highly subjective and inaccurate, valid diagnosis requires prospectively evaluating daily symptoms against the diagnostic criteria<sup>3,4</sup>. Typically, symptoms are evaluated through visual inspection of daily ratings across multiple cycles; however, differences in interpretation of the diagnostic criteria during these visual inspections may lead to inconsistencies. Furthermore, the complex diagnostic criteria outlined in the DSM-5 means there is a high risk of diagnostic error due to differing practices and errors in clinical judgment. These concerns suggest the need for a more reliable method for making valid prospective diagnoses. In response to this need, Eisenlohr-Moul et al. (under review) have developed the Carolina Premenstrual Assessment Scoring System (or C-PASS), a computerized standardized

diagnostic system aimed at creating shared meaning for PMDD and MRMD across laboratories. Although the C-PASS, described in more detail below, represents an essential step forward in our standardization of these diagnoses, additional work is needed to verify that the cut points selected for making the diagnosis are adequate for identifying the subgroup of women that need treatment. The purpose of the present paper is to use descriptive statistics and ROC analyses to examine how changes to these diagnostic cutoffs influence the frequency and appropriateness of diagnoses.

# Defining the DSM-5 Diagnostic Dimensions and Mapping Them onto a Widely-Used Assessment Tool

The DSM-5 outlines four key diagnostic dimensions for characterizing PMDD: content, cyclicity, clinical significance, and chronicity. The <u>content</u> dimension requires that five symptoms are met and at least one of those symptoms must be a core symptom such as depression, anxiety, mood lability, or anger/irritability. A MRMD diagnosis only requires that one core symptom is present. The <u>cyclicity</u> dimension refers to the relative symptom elevation and the absolute symptom clearance. Relative premenstrual symptom elevation is met if the symptom shows at least a 30% decrease in severity from the premenstrual week to the postmenstrual week. Absolute symptom clearance is met if a symptom does not exceed a Likert score of three out of a maximum of six on any day during the postmenstrual week. The <u>clinical significance</u> dimension refers to the absolute severity of the symptoms and their duration. This dimension requires that the symptoms must be of sufficient absolute severity and duration in the pre-menstrual week to be classified as clinically significant. The

<u>chronicity</u> dimension requires that the dimensions of content, cyclicity, and clinical significance be present for a majority of the cycles.

The DSM-5 does not provide numerical equivalences for the diagnostic dimensions, which is part of the reason for subjectivity and variability in PMDD diagnostic practices. However, the Daily Record of Severity Problems form (DRSP) allows for the numerical assessment of the criterion described for the four diagnostic dimensions (Table 9). The DSM-5 content for PMDD and MRMD is completely contained within the DRSP's twenty-one symptoms. The DRSP asks the rater to indicate the daily severity of these twenty-one symptoms on a 6-point Likert scale. A score of one indicates the symptom was not at all present and a score of six indicates an extreme presence of the symptom.

The C-PASS was created to standardize the complicated PMDD diagnosis. While the diagnosis in the C-PASS was created according to the DSM-5, certain numeric threshold decisions were made to ensure that the C-PASS outcome was reasonable and warranted diagnosis and treatment. Although the C-PASS represents an important first step in standardizing the diagnoses of PMDD and MRMD, the present paper examines whether the cutoff values on the DRSP that were selected for the C-PASS criteria are appropriate in terms of selecting a group of women that requires treatment. In this paper, we re-examine the sample of women seeking a diagnosis of MRMD, using descriptive statistics and logistic/ROC analyses to understand whether the current cut points are appropriate for identifying women in need of treatment.

Several variations in calculation methods and thresholds for diagnosis were considered. First, we compare diagnostic frequencies using follicular maximum ratings

or follicular mean ratings to evaluate absolute clearance (see above). Second, we assessed descriptive changes when altering the thresholds for the number of symptoms needed per cycle, relative symptom elevation threshold, severity and clearance threshold, and the threshold for duration of severe symptoms. In order to determine the number of symptoms per cycle and the number of severe days (duration) per cycle at which cyclicity of life impairment is present, we also conducted an ROC analyses for these predictors (using the efficiency criterion to select an optimal cutpoint). These analyses will be described and discussed in terms of their relevance for understanding the disorder and improving the C-PASS diagnostic system.

#### Methods

#### **Description of the C-PASS Diagnostic Method**

The diagnostic process begins by characterizing each DRSP item in each cycle (where a cycle is defined as a set of contiguous postmenstrual and premenstrual weeks from two consecutive menstrual cycles) using the **four diagnostic dimensions** as described in Table 10 (*relative symptom elevation:* percent symptom elevation during premenstrual phase relative to postmenstrual phase >=30%; *absolute clearance:* postmenstrual week maximum <=3; *absolute severity:* premenstrual week maximum >=4; and *duration:* severe premenstrual week days >=2). Because DSM-5 diagnosis of PMDD is clearly defined as a marked on-off pattern occurring in the *perimenstrual timeframe*, the C-PASS utilizes the premenstrual week (defined as days -7 to -1, where -1 is the day prior to menstrual onset) and the postmenstrual week (defined as the 7 days following average menstrual offset: days 4 to 10, where day 1 represents

menstrual onset). That is, the rationale for comparing the premenstrual week of one menstrual cycle to the contiguous follicular postmenstrual week of the next cycle is to establish the "switch off" of symptoms, as it is critical to demonstrate that the cyclical symptoms do not persist into the follicular phase. Further, the C-PASS requires that at least 3 out of 7 ratings be present in each of the two weeks from each cycle, and requires at least two cycles (i.e., contiguous pairs of premenstrual-postmenstrual phases).

Next, cycle-level diagnosis of PMDD is made by counting DSM-5 symptoms meeting criteria on all four dimensions (see Table 9; Total Symptoms: 1-4 for MRMD and >=5 for PMDD) and whether a core symptom meets criteria (number of core symptoms >=1). Next, the C-PASS makes the diagnosis of MRMD or PMDD at the person level by counting the number of cycles meeting diagnostic criteria for either MRMD or PMDD (cycles meeting criteria >=2). Finally, the C-PASS concatenates a visual representation of relevant information for each DRSP item across as many cycles as provided, along with a determination of cycle-level diagnosis for that symptom in each cycle. The system also outputs a dataset with individual difference variables for each diagnostic dimension. In the present study, the research diagnosis of MRMD<sup>5</sup> (i.e., 1-4 symptoms met for at least 2 cycles, of which one must be a core emotional symptom) was calculated in addition to PMDD.

#### Participants, Procedure, and Materials

The study sample consisted of 267 naturally cycling women between ages 18 and 47 (M = 32.70, SD = 8.21) with regular menstrual cycles (21–35 days). Women with

pregnancy, chronic medical disorders, history of certain psychiatric symptoms (mania, substance dependence, psychosis), or certain prescription medications (any antidepressant, benzodiazepines, neuroleptics, or any hormonal preparation) were excluded. All procedures were approved by the IRB, and all participants provided informed consent. These participants were not paid.

Subjects were recruited at the University of North Carolina using posters, flyers, and e-mail over a six-year period (2009-2015). All recruitment materials specified that the purpose of this study was to assess menstrual cycle-related psychological and somatic problems. An initial telephone screening was conducted to assess inclusion and exclusion criteria. Baseline visits were also scheduled for eligible participants during these interviews. During the 45-minute baseline visits, subjects' medical and medication histories were recorded, and the SCID-1 was administered to assess for Axis I psychiatric disorders.

Prospective assessment of menstrual cycle-related symptoms was completed using 2-4 months of daily participant diaries. Participants completed daily ratings using the DRSP (see above for description) and reported menstrual flow. Participants were also able to report external events, circumstances, or stressors they believe may be associated with their daily well-being. Days where participants reported the occurrence of a substantial external stressor were treated as missing data. These forms were mailed in weekly. In the final sample, 200 women had provided sufficient prospective data and 67 women had been eliminated based upon an insufficient amount of data.

#### **Analytic Plan**

Descriptive tables were produced to compare the use of two different statistics for the evaluation of absolute premenstrual severity and absolute postmenstrual symptom clearance: follicular maximum rating versus follicular mean rating. Four different thresholds were then varied for relative symptom elevation, absolute severity and clearance, duration, number of severe symptoms. After conducting multilevel (cycles nested within women) logistic regressions, receiver operating curves were created to determine the optimal cutpoint for both 1) number of symptoms per cycle and 2) number of severe days per cycle on each DRSP item. The dichotomous outcome was whether or not the cycle met C-PASS criteria on any impairment item. Impairment is met when a symptom meets C-PASS criteria for any of the following three DRSP items: interference with work, interference with hobbies or social life, and interference with relationships. The symptom must meet C-PASS criteria for at least one of C-PASS criteria for cyclical impairment to be met. SAS 9.5 ROCPLOT macro was used to determine the optimal cut points using the efficiency method. The efficiency calculation utilizes prevalence to weight the specificity and sensitivity and is calculated using the following formula:

While other optimal cut point calculations in SAS 9.3, such as the Youden index, assume a 50% base rate within the sample, the efficiency method takes the true base rate of the sample into account. An assumption of a 50% base rate was not deemed appropriate given the base rates described earlier. Using the base rates within the sample will provide us with estimates most accurate for women seeking PMDD and MRMD diagnoses. Furthermore, we opted for a cut point calculation method that does

not account for the cost of incorrect diagnosis. Currently, there is little evidence available on which to conduct a cost-benefit analysis regarding the clinical detriments of false positives vs. false negatives in PMDD; thus the decision was made to ignore the cost ratio dimension.

#### Results

#### **Preliminary Analyses**

267 women contributed 563 cycles to the analysis. 170 cycles (30.20%) were characterized by significant cyclical impairment, 285 cycles (50.62%) received the C-PASS MRMD diagnosis, and 149 cycles (26.46%) received the more stringent PMDD diagnosis. All analyses were carried out in SAS 9.5. Each cycle also received a dichotomous decision regarding the presence of significant cyclical interference following the C-PASS method for diagnosing symptoms described above. Relative premenstrual symptom elevation may be calculated using either the phase average severity for each symptom or the phase maximum severity for each symptom. Diagnosis frequencies within the sample were compared when each method of calculation was used (Table 1). Upon inspection of the descriptive statistics, it appears that using follicular mean in the calculation is a less stringent method of diagnosis. Further, within each method of absolute symptom clearance calculation (follicular maximum and follicular mean), the following thresholds were varied and descriptives presented: percent threshold for relative premenstrual symptom elevation, the absolute severity and clearance threshold, the symptom duration thresholds, and the number of symptoms required to meet criteria were varied.

Table 1. Comparing Changes in C-PASS Diagnosis when varying premenstrual symptom elevation calculation

	C-PASS Diagnosis with Relative Premenstrual Symptom Elevation Calculation Varied					
•	No Diagnosis	MRMD Diagnosis	PMDD Diagnosis			
Calculation Method	FREQ	FREQ	FREQ			
1) Using Follicular Mean to						
Evaluate Diagnostic	<b>103</b> (53%)	<b>48</b> (25%)	<b>43</b>			
Thresholds	(55%)	(23%)	(22%)			
2) Follicular Max to Evaluate	114	45	35			
Diagnostic Thresholds	(59%)	(23%)	(18%)			

Relative Symptom Elevation Thresholds. Relative symptom elevation calculation using mean severity for the follicular phase resulted in 22% of the participants diagnosed with PMDD when a 30% threshold for relative symptom elevation was specified (Table 2). This number of patients diagnosed decreased dramatically when the threshold for relative symptom elevation was increased to 50% (10% diagnosed with PMDD). Trends were similar when follicular maximum was used.

Table 2. Impact of Varying % Threshold for Relative Premenstrual Symptom Elevation

	C-P/	C-PASS Diagnosis with Relative Symptom Elevation Varied					
	No Dia	No Diagnosis		MRMD Diagnosis		Diagnosis	
Throchold	Using Fol	Using Fol	Using Fol	Using Fol	Using Fol	Using Fol	
<u>Threshold</u>	Mean	Max	Mean	Max	Mean	Max	
1) <b>\( \Delta \)</b> 30%	<b>103</b> (53%)	<b>114</b> (59%)	<b>48</b> (25%)	<b>45</b> (23%)	<b>43</b> (22%)	<b>35</b> (18%)	
2) Δ 50%	<b>156</b> (80%)	<b>161</b> (83%)	<b>18</b> (9%)	<b>19</b> (10%)	<b>20</b> (10%)	<b>14</b> (7%)	
3) ∆ 75%	<b>187</b> (96%)	<b>187</b> (96%)	<b>5</b> (3%)	<b>5</b> (3%)	<b>2</b> (1%)	<b>2</b> (1%)	

Absolute Symptom Severity and Clearance Thresholds. Absolute severity thresholds greater than or equal to 3 and absolute clearance thresholds less than or equal to three resulted in a 28% diagnosis of PMDD (Table 3). Although this percentage on PMDD diagnosis decreased (22%) when the threshold was increased to 4, the percentage of MRMD diagnoses remained the same (25%). A threshold of 5 further decreased the percentage of MRMD diagnosis and PMDD diagnosis (Table 3). Trends were similar when follicular maximum was used.

Table 3. Determination of Absolute Severity and Clearance Thresholds

	C-PASS Diagnosis with Absolute Severity and Clearance Varied						
	No Dia	agnosis	MRMD D	MRMD Diagnosis		Diagnosis	
Threshold	Using Fol Mean	Using Fol Max	Using Fol Mean	Using Fol Max	Using Fol Mean	Using Fol Max	
<ol> <li>Abs Severity &gt; 3</li> <li>Abs Clearance ≤ 3</li> </ol>	<b>91</b> (47%)	<b>103</b> (52%)	<b>49</b> (25%)	<b>48</b> (24%)	<b>54</b> (28%)	<b>49</b> (19%)	
2) Abs Severity > 4 Abs Clearance ≤ 4	<b>103</b> (53%)	<b>114</b> (59%)	<b>48</b> (25%)	<b>45</b> (23%)	<b>43</b> (22%)	<b>35</b> (18%)	
3) Abs Severity > 5 Abs Clearance ≤ 5	<b>132</b> (68%)	<b>134</b> (74%)	<b>32</b> (16%)	<b>28</b> (15%)	<b>30</b> (15%)	<b>19</b> (11%)	

**Duration Thresholds.** The duration of symptoms was varied from two to five days and frequencies of PMDD and MRMD diagnosis generally decreased as the threshold of days increased (Table 4). Trends were similar when follicular maximum was used.

Table 4. Duration of Symptoms

	C-PASS Diagnosis with Duration Varied					
	No Dia	No Diagnosis		MRMD Diagnosis		Diagnosis
<u>Threshold</u>	Using Fol Mean	Using Fol Max	Using Fol Mean	Using Fol Max	Using Fol Mean	Using Fol Max
1) ≥ 2 Days	<b>103</b> (53%)	<b>114</b> (59%)	<b>48</b> (25%)	<b>45</b> (23%)	<b>43</b> (22%)	<b>35</b> (18%)
2) ≥3 Days	<b>120</b> (62%)	<b>134</b> (69%)	<b>38</b> (20%)	<b>33</b> (17%)	<b>36</b> (19%)	<b>27</b> (14%)
3) ≥4 Days	<b>140</b> (72%)	<b>153</b> (79%)	<b>27</b> (14%)	<b>24</b> (12%)	<b>27</b> (14%)	<b>17</b> (9%)

**Symptom Thresholds.** The number of severe symptoms needed for each cycle to meet C-PASS criteria was also varied. Table 5 shows the frequency results as number of symptom threshold was varied from two to five. The current threshold is set at five symptoms. As one would expect, as the stringency of the threshold is increased, the frequency of diagnosis decreases.

Table 5. Number of Symptoms meeting criteria to diagnose PMDD

	C-PASS Diagnosis with No. of Total Symptoms Varied				
•	No Dia	ignosis	PMDD Diagnosis		
Calculation Method	Using Fol Mean	Using Fol Max	Using Fol Mean	Using Fol Max	
1) # Total SX ≥ 2	<b>114</b> (59%)	<b>124</b> (64%)	<b>80</b> (41%)	<b>70</b> (36%)	
2) # Total SX ≥ 3	<b>126</b> (65%)	<b>135</b> (70%)	<b>68</b> (35%)	<b>59</b> (30%)	
3) # Total SX ≥ 4	<b>140</b> (72%)	<b>147</b> (76%)	<b>54</b> (28%)	<b>47</b> (24%)	
4) # Total SX ≥ 5	<b>151</b> (78%)	<b>159</b> (82%)	<b>43</b> (22%)	<b>35</b> (18%)	

Using Multilevel Logistic Regression and ROC Curves to Determine the Optimal Number of 1) DSM-5 Symptoms per Cycle and 2) Severe Days per Cycle to Predict the Presence of Moderate Cyclical Impairment

Next logistic multilevel models in which cycles were nested within women were used to predict cycle-level presence of significant cyclical interference from number of DSM-5 symptoms meeting C-PASS criteria in that cycle; number of symptoms meeting criteria was indeed predictive of whether or not the cycle was characterized by impairment (*Estimate*: .59, SE = .049, t(295) = 11.99, p < .0001). Random effects for number of symptoms were not significant, suggesting that the size and direction of the effect of number of symptoms per cycle on impairment in a given cycle was similar across women. Information regarding predicted values was saved from this model and averaged across each number of symptoms. PROC LOGISTIC was utilized to calculate the area under the ROC curve (AUC = .9017, 95% CI: .87 to .92). Finally, the SAS ROCPLOT macro<sup>6</sup> was utilized to graph the ROC curve (see Figure 1) and the efficiency criterion was utilized to select the optimal number of symptoms per cycle for predicting the presence of significant cyclical interference, which was 4 symptoms.

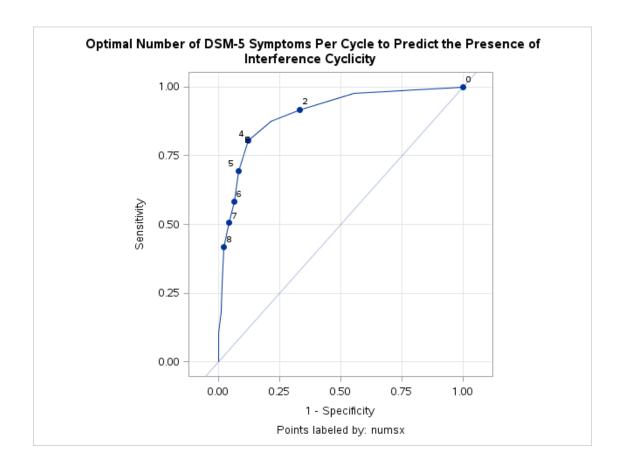


Figure 1. ROC Curve Predicting Presence of Impairment in this Cycle from Number of Symptoms Meeting DSM-5 Criteria this Cycle

Table 6. ROC Curve Analysis Results for Number of Symptoms Meeting DSM-5 Criteria Predicting the Presence of Impairment

AUROC	Confider	nce Limit	Cutpoint*	Optimal Number of Symptoms
0.9017	0.8742	0.9293	-0.41359	4

Therefore, despite the fact that the DSM-5 does not require the presence of interference to make the diagnosis of PMDD (this is a shift from DSM-IV), this ROC analysis suggests that the number of DSM-5 symptoms per cycle at which significant cyclical impairment can be optimally predicted is 4, which is LOWER than the number of symptoms per cycle (5) needed to make the official diagnosis of DSM-PMDD. In similar ROC analyses conducted for the number of severe days on each DRSP item needed to predict moderate cyclical impairment, the number of severe days needed to cause impairment by each symptom ranged from two days to seven days. In general, it appears that high arousal symptoms such as anxiety and irritability require a greater amount of severe days within a cycle to cause impairment. Low arousal psychological symptoms require a lower number of severe days to cause impairment. Most DRSP items required either three days or five days to cause impairment. Physical symptoms generally needed five to seven days to cause impairment, indicating that they are may not be the key symptoms of PMDD in terms of driving impairment. A summary of these ROC analyses can be found in Tables 7 and 8. Table 8 excludes DRSP 2, headaches, because this is not considered a symptom indicative of PMDD.

Table 7. Summary of ROC curve analyses results for number of severe days predicting the presence of at least moderate impairment cyclicity.

Predictor	AUROC	Confider	nce Limit	Cutpoint*	Optimal Number of Days
DRSP 1	0.7425	0.6974	0.7877	0.22176	4
DRSP 2	0.6839	0.6388	0.7290	-0.41264	2
DRSP 3	0.6796	0.6357	0.7234	-0.063947	3
DRSP 4	0.7649	0.7217	0.8080	0.23155	5
DRSP 5	0.7868	0.7457	0.8279	0.40310	5
DRSP 6	0.7870	0.7463	0.8278	-0.25939	3
DRSP 7	0.8345	0.7995	0.8694	0.53480	5
DRSP 8	0.8185	0.7803	0.8567	-0.096879	3
DRSP 9	0.7919	0.7510	0.8328	-0.097786	3
DRSP 10	0.7741	0.7318	0.8164	-0.17985	3
DRSP 11	0.7872	0.7465	0.8279	0.33197	5
DRSP 12	0.7391	0.6943	0.7839	-0.29633	3
DRSP 13	0.7181	0.6722	0.7640	0.070238	4
DRSP 14	0.7565	0.7137	0.7994	0.029269	4
DRSP 15	0.6927	0.6445	0.7409	0.52389	5
DRSP 16	0.7620	0.7185	0.8054	-0.057696	3
DRSP 17	0.7650	0.7225	0.8075	0.20529	3
DRSP 18	0.6735	0.6268	0.7202	0.10991	5
DRSP 19	0.7314	0.6868	0.7760	0.87214	7
DRSP 21	0.6572	0.6098	0.7045	0.53208	5

Table 8. The number of severe days required on each DRSP item to optimally predict the presence of at least moderate impairment cyclicity.

2 Days	3 Days	4 Days	5 Days	7 Days
Hopelessness (2)	Worthlessness (3)	Depressed/Blue (1)	Anxious (4)	Breast Swelling/Bloated (19)
Headache (20)	Sensitivity to Rejection (6)	Specific Food Cravings (13)	Mood Swings (5)	(10)
	Conflicts with People (8)	Slept More (14)	Angry/Irritable (7)	
	Less interest in usual activities (9)		Lethargic/ Lack of Energy (11)	
	Difficulty Concentrating (10)		Trouble Sleeping (15)	
	Increased Appetite (12)		Breast Tenderness (18)	
	Overwhelmed (16)		Joint/Muscle Pain (21)	
	Feeling Out of Control (17)			

#### **Discussion**

Variations in approaches to making the diagnosis of premenstrual dysphoric disorder compromise the validity of PMDD and undermine our ability to identify and treat women suffering from the disorder. The Carolina Premenstrual Assessment Scoring System (C-PASS) seeks to streamline the diagnosis process and create diagnostic uniformity within the field. Although many of the diagnostic decisions made within the C-PASS have been recommended by the DSM-5, certain decisions made within the C-PASS required further inspection to understand how changing them might alter the

prevalence of diagnosis. The sample used consisted of women seeking evaluation for PMDD and willing to participate in a research study.

Descriptive statistics were calculated for a variety of permutations of thresholds and methods to provide information about how changing these thresholds and methods might influence diagnostic prevalence. ROC curves were examined regarding the number of symptoms per cycle needed to predict moderate impairment, which is an outcome used in order to identify a subgroup of women who require treatment. ROC curves were also created for each DRSP symptom to determine the optimal number of severe days required to see at least moderate impairment.

# **Findings and Implications**

The ROC in figure 1 shows that four symptoms is the optimal number of symptoms that should be required to diagnose PMDD. This finding is consistent with findings from the Harvard Study of Moods and Cycles, which found that a community-based sample needed six symptoms while a clinical sample only required two symptoms<sup>7</sup>. The average value of these two very different samples once again confirms that four symptoms is likely the ideal number for diagnosing PMDD. This threshold of four symptoms differs from the five symptom requirement by the DSM-5. Our findings suggest that the current five symptom threshold in the DSM-5 may be too stringent and cause missed diagnoses.

In general the ROC curves (Tables 6 and 7) showed that 3 days and 5 days of severity were sufficient to cause impairment by most symptoms. Table 7 shows the vast differences in each DRSP item and the optimal number of days to cause cyclical

impairment by that item. Therefore, the current DSM-5 requirement of 2 severe days per symptom might potentially be too lax.

Variations of threshold produced the expected effect of diagnosing more patients as the thresholds were made more stringent. It does not appear that utilizing the follicular maximum versus the follicular mean creates a large, substantial difference in the number of PMDD diagnoses made. The C-PASS utilizes follicular maximum, and this decision appears to be justified at the present time.

## **Limitations of this study**

While this study examined impairment caused by PMDD symptoms, the DRSP interference items were the only measures used to establish impairment. Further work could be improved by including alternative validity measures at the daily level for both distress and impairment. Furthermore, this study has issues of generalizability. The women included in this study were actively seeking aid for PMDD and were willing to participate in research studies. This may not be representative of the entire community of women suffering from PMDD, and it is likely that many "silent sufferers" of the disease may not have been represented. A community-based cohort would provide a more representative group. On the other hand, this sample does represent the current normative sample used by most laboratories when studying PMDD—women who respond to advertisements about premenstrual symptoms.

### **Conclusions**

The Carolina Premenstrual Assessment Scoring System seeks to systematically diagnose and standardize premenstrual dysphoric disorder. The descriptive examination of diagnostic thresholds presented here demonstrates that, while the majority of the decision rules incorporated in the C-PASS are defensible, further work is needed to determine whether some of the decision rules should be altered. In particular, the current threshold of five symptoms to reach PMDD diagnosis may be too stringent, given that just four symptoms were necessary in the present study to predict impairment cyclicity, and the current threshold of two severe premenstrual days of symptoms may be too lax, given that four severe days were necessary here in order to predict impairment cyclicity.

# Table 9. Mapping the Items of the DRSP onto DSM-5 Diagnostic Content

DRSP	DSM-5		
CORE SYMPTON	IS / CRITERION B		
DRSP 5. Had mood swings (e.g. suddenly felt sad or tearful)	Marked affective lability (e.g., mood swings; feeling		
DRSP 6. Was more sensitive to rejection or my feelings were easily hurt	suddenly sad or tearful, or increased sensitivity to rejection)		
DRSP 7. Felt angry, irritable	Marked irritability or anger or increased interpersonal		
DRSP 8. Had conflicts or problems with people	conflicts		
DRSP 1. Felt depressed, sad, "down" or blue			
DRSP 2. Felt hopeless	Marked <b>depressed mood,</b> feelings of <b>hopelessness</b> , or self-deprecating thoughts		
DRSP 3. Felt worthless or guilty			
DRSP 4. Felt anxious, "keyed up", or "on edge"	Marked <b>anxiety</b> , <b>tension</b> , and/or feelings of being keyed up or on edge		
ADDITIONAL SYMP	TOMS / CRITERION C		
DRSP 9. Had less interest in usual activities (e.g. work, school, friends, hobbies)	Decreased interest in usual activities (e.g. work, school, friends, hobbies)		
DRSP 10. Had difficulty concentrating	2. Subjective difficulty in concentration		
DRSP 11. Felt lethargic tired, fatigued, or had a lack of energy	3. Lethargy, easy fatigability, or marked lack of energy		
DRSP 12. Had increased appetite or overate	Marked change in appetite; overeating; or specific food		
DRSP 13. Had specific food cravings	cravings		
DRSP 14. Slept more, took naps, found it hard to get up			
DRSP 15. Had trouble getting to sleep, staying asleep	5. Hypersomnia or Insomnia		
DRSP 16. Felt overwhelmed, that I couldn't cope	6. A copies of being everybalmed or out of control		
DRSP 17. Felt out of control	A sense of being overwhelmed or out of control		
DRSP 18. Had breast tenderness			
DRSP 19. Had breast swelling, felt bloated, or had weight gain	7. Physical symptoms such as breast tenderness or swelling, joint or muscle pain, sensation of "bloating", or weight gain		
DRSP 21. Had Joint or muscle pain			
DRSP 20. Had headache			

Table 10. Diagnostic Dimensions of DSM-5 Premenstrual Dysphoric Disorder

DIAGNOSTIC	C DIMENSIONS	Diagnosis	Based on DRSP	DSM-5
Content	Symptoms	with other people Secondary symptoms: less interest in usual activities, difficu lethargic/fatigue/tired/lack of energy, cravings, slept more/took naps/hard t	d swings, was more sensitive to lt angry/irritable, had conflicts/problems alty concentrating, increased appetite/overate, specific food to get up, trouble getting to sleep/staying pe, felt out of control, breast tenderness, iin, headache, joint or muscle pain ome or in daily routine"	Criterion B: affective lability, irritability/anger/increased interpersonal conflicts, depressed mood/feelings of hopelessness/self-deprecating thoughts, anxiety/tension/feelings of being keyed up/on edge Criterion C: decreased interest, difficulty in concentration, lethargy/easy fatigability/lack of energy, change in appetite, hypersomnia/insomnia, overwhelmed/out of control, physical symptoms (breast tenderness, muscle pain, bloating, weight gain)
	Number	MRMD ≥ 1 core symptom	PMDD ≥ 1 core symptom ≥ 5 total symptoms	Criterion A: A total of 5 [at least (one or more) of each subgroup]
0.11.11	Relative Premenstrual Elevation	menstrual onse and 1 is menstrual onset  Symptoms must not exceed a value of 3 on any day during days 4 → 10		Criterion A: "present in the week before mensesimprove within a few days after the onset of menses"
Cyclicity	Absolute Postmenstrual Clearance			Criterion A:  "minimal or absent in the week postmenses"  Postmenses = following menstrual onset
Clinical	Absolute Premenstrual Severity	4 or more (on a Likert-scale from 1 to	0 6)	Criterion D:  "symptoms are associated with clinically significant distress  OR interference with work, school, usual social activities, or relationships with others"
Significance	Premenstrual Duration	At least 2 days (doesn't have to be consecutive)		Criterion D: "in the final week before the onset of menses"
Not Simply Cyclicity of Other Disorder		Rule out dysmenorrhea using prospective ratings. Rule out mood and anxiety disorder with SCID-1. Rule out Borderline Personality Disorder with SCID-2.		Criterion E:  "not merely an exacerbation of the symptoms of another disorder."  "Key differential diagnoses: dysmenorrhea, bipolar disorder, MDD, dysthymia, and BPD."
≥ 2 symptomatic months  Chronicity			Criterion A and F: "In the majority of menstrual cycles" "should be confirmed by prospective daily ratings during at least two symptomatic cycles."	

#### References

- 1. Pearlstein T, Steiner M. Premenstrual dysphoric disorder: burden of illness and treatment update. J Psychiatry Neurosci. 2008;33(4):291-301.
- 2. Halbreich U, Borenstein J, Pearlstein T, Kahn LS. The prevalence, impairment, impact, and burden of premenstrual dysphoric disorder (PMS/PMDD). *Psychoneuroendocrinology*. 2003;28:1-23. doi:10.1016/S0306-4530(03)00098-2.
- 3. Smith MJ, Schmidt PJ, Rubinow DR. Operationalizing DSM-IV criteria for PMDD: selecting symptomatic and asymptomatic cycles for research. *Journal of Psychiatric Research*. 2003;37(1):75-83.
- 4. Hartlage SA, Freels S, Gotman N, Yonkers K. Criteria for Premenstrual Dysphoric Disorder: Secondary Analyses of Relevant Data Sets. Arch Gen
- 5. Dean BB, Borenstein JE, Knight K, Yonkers K. Evaluating the Criteria Used for Identification of PMS. http://dxdoiorg/101089/jwh200615546. 2006;15(5):546-555. doi:10.1089/jwh.2006.15.546.
- 6. 25018 Plot ROC curve with cutpoint labeling and optimal cutpoint analysis. (n.d.). Retrieved March 10, 2016, from <a href="http://support.sas.com/kb/25/018.html">http://support.sas.com/kb/25/018.html</a>
- 7. Cohen LS, Soares CN, Otto MW, Sweeney BH, Liberman RF, Harlow BL. Prevalence and predictors of premenstrual dysphoric disorder (PMDD) in older premenopausal women The Harvard Study of Moods and Cycles. *Journal of Affective Disorders*. 2002; 70(2):125-132.