# Depression and Anxiety in Persons with Von Willebrand Disease

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#### Introduction

- Depression and anxiety are associated with poor health-related quality of life (HRQoL), lower functioning and decreased treatment adherence
- In 2019, 7% of adults in the US had moderate/severe symptoms of depression, while <5% had anxiety based on 2019 data of National Center for Health Statistics
- Impacts of depression and anxiety in persons with von Willebrand disease (VWD) are unclear
- Objective: To assess sociodemographic and clinical characteristics associated with depression and anxiety in a geographically diverse cohort of individuals with VWD obtaining care at seven US Hemophilia Treatment Centers (HTCs)

### Methods



- □ Study participants: age  $\geq$ 12 with VWD Type 1 (VWF:Ag/RCo:  $\leq$ 30%), low VWF (VWF:Ag/RCo: 30-50%), Type 2, and 3
- Survey collected sociodemographic and clinical data, self-reported pain, joint problems and HRQoL measured by EQ-5D-3L at enrollment, as well as quarterly follow-up for one year
- PHQ-8 and GAD-7 were administered with the last follow-up survey after August 2019
  - Score values ≥10 on either PHQ-8 or GAD-7 were classified as depression or anxiety, respectively
- The association of sociodemographic and clinical characteristics with depression or anxiety was assessed using Chi-square tests for categorical variables, as well as logistic regression models with stepwise selection

#### Results: Participants Characteristics by Depression or Anxiety



|  |                 | Depression         |                   |             | Anxiety         |                    |                   |             |
|--|-----------------|--------------------|-------------------|-------------|-----------------|--------------------|-------------------|-------------|
| Variable   | Total<br>(N=77) | Yes<br>(N=49, 64%) | No<br>(N=28, 36%) | P<br>Value* | Total<br>(N=72) | Yes<br>(N=42, 58%) | No<br>(N=30, 42%) | P<br>Value* |
| Mean (SD) age in years                             | 34.2 (18.8)     | 31.0 (17.8)        | 39.8 (19.4)       | 0.047       | 33.5 (18.4)     | 27.0 (13.9)        | 42.7 (20.2)       | 0.001       |
| Low VWD  |                 |                    |                   | 0.04        |                 |                    |                   | 0.38        |
| Type 1, 2, 3                                       | 62 (80.5)       | 36 (58.1)          | 26 (41.9)         |             | 59 (81.9)       | 33 (55.9)          | 26 (44.1)         |             |
| Low VWF  | 15 (19.5)       | 13 (86.7)          | 2 (13.3)          |             | 13 (18.1)       | 9 (69.2)           | 4 (30.8)          |             |
| Gender   |                 |                    |                   | 0.92        |                 |                    |                   | 0.30        |
| Male   | 16 (20.8)       | 10 (62.5)          | 6 (37.5)          |             | 15 (20.8)       | 7 (46.7)           | 8 (53.3)          |             |
| Female   | 61 (79.2)       | 39 (63.9)          | 22 (36.1)         |             | 57 (79.2)       | 35 (61.4)          | 22 (38.6)         |             |
| Marital status                                     |                 |                    |                   | < 0.01      |                 |                    |                   | < 0.01      |
| Married/with a partner                             | 53 (70.7)       | 28 (52.8)          | 25 (47.2)         |             | 50 (71.4)       | 24 (48.0)          | 26 (52.0)         |             |
| Single/not with a partner                          | 22 (29.3)       | 19 (86.4)          | 3 (13.6)          |             | 20 (28.6)       | 17 (85.0)          | 3 (15.0)          |             |
| General health compared to 3 months ago            |                 |                    |                   | 0.02        |                 |                    |                   | 0.05        |
| Better/same  | 64 (83.1)       | 37 (57.8)          | 27 (42.2)         |             | 60 (83.3)       | 32 (53.3)          | 28 (46.7)         |             |
| Worse  | 13 (16.9)       | 12 (92.3)          | 1 (7.7)           |             | 12 (16.7)       | 10 (83.3)          | 2 (16.7)          |             |
| Self-reported chronic pain                         |                 |                    |                   | 0.0003      |                 |                    |                   | 0.62        |
| No pain  | 19 (27.9)       | 7 (36.8)           | 12 (63.2)         |             | 19 (29.7)       | 11 (57.9)          | 8 (42.1)          |             |
| Any pain   | 49 (72.1)       | 40 (81.6)          | 9 (18.4)          |             | 45 (70.3)       | 29 (64.4)          | 16 (35.6)         |             |
| Self-reported joint problems                       |                 |                    |                   | 0.002       |                 |                    |                   | 0.02        |
| Yes  | 34 (44.2)       | 28 (82.4)          | 6 (17.6)          |             | 31 (43.1)       | 23 (74.2)          | 8 (25.8)          |             |
| No   | 43 (55.8)       | 21 (48.8)          | 22 (51.2)         |             | 41 (56.9)       | 19 (46.3)          | 22 (53.7)         |             |
| EQ-VAS adjusted mean (SE) <sup>+</sup>             |                 | 68.77 (3.15)       | 77.58 (4.24)      | 0.03        |                 | 69.27 (3.72)       | 71.29 (4.25)      | 0.66        |
| EQ-5D index score adjusted mean (SE) <sup>+</sup>  |                 | 0.75 (0.03)        | 0.83 (0.04)       | 0.06        |                 | 0.75 (0.03)        | 0.82 (0.04)       | 0.07        |
| Chronic pain score adjusted mean (SE) <sup>+</sup> |                 | 3.65 (0.45)        | 2.50 (0.65)       | 0.08        |                 | 3.20 (0.48)        | 2.92 (0.59)       | 0.67        |

Abbreviations: SD, standard deviation; VWD, Von Willebrand disease; SE, standard error; EQ-VAS, EuroQoL visual analogue scale. Note: data were presented as frequency (row percentage) for categorical variables or mean (standard deviation) for continuous variables. \*P values were calculated from Chi-square tests for categorical variables and T-tests for continuous variables. †Covariates in included age, gender, employment, VWD type, self-reported joint problem, self-rate general health compared to 3 month ago.



#### Results: Predictors of Depression or Anxiety

|  | Standardized         | Odds Ratio          |         |       |
|--|----------------------|---------------------|---------|-------|
| Variables  | <b>β</b> Coefficient | (95% CI)            | P value | Step* |
| Depression (N=74)  |                      |                     |         |       |
| Self-reported joint problems (Yes vs No) <sup>+</sup>                | 0.51                 | 6.3 (1.99-20.09)    | 0.002   | 1     |
| Marital status: Single/not with a partner vs Married/with a partner  | 0.49                 | 7.0 (1.69-29.03)    | 0.007   | 2     |
|  |                      |                     |         |       |
| Anxiety (N=69)   |                      |                     |         |       |
| Marital status (Single/not with a partner vs Married/with a partner) | 0.60                 | 10.80 (2.46-47.46)  | 0.002   | 1     |
| Age (12-17 vs ≥18 years old)   | 0.45                 | 6.66 (1.65-26.92)   | 0.008   | 2     |
| Health compared to 3 months prior (Worse vs Better/same)             | 0.51                 | 12.34 (1.31-116.21) | 0.03    | 3     |

Abbreviations: CI, confidence interval; N, indicates sample size.

Note: Participants with missing data were excluded from the regression analyses. The variables which were included in the model included age, gender, marital status, employment, VWD type, self-reported joint problems, and health compared to 3 months prior.

\*The order of the variable entered into the multiple logistic regression model.

<sup>+</sup>Self-reported joint problems were assessed by the question "Do you have any problems with your joints?".



#### Conclusions

- Our study revealed higher rates of major depression and anxiety in this VWD sample than the general US population
- Depression had a significant negative impact on HRQoL
- Mental health screening is critical for persons with VWD, especially those with low VWF, chronic pain or joint problems
- Special attention should be paid to women and youth
- This study underscores the need for optimal mental health service as part of the multidisciplinary approach in the comprehensive care of persons with VWD seen at HTCs

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### Conflict of Interest

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