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Depression, anxiety and perceived stress in women with and without PCOS: a community-based study

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Abstract

Background. Polycystic ovary syndrome (PCOS) is associated with increased psychological distress in clinical populations. We aimed to assess depression, anxiety and perceived stress in women with and without PCOS in a large community-based sample and investigate the role of stress in contributing to and mediating the relationship between PCOS, depression and anxiety.

Methods. A cross-sectional analysis was performed from the Australian Longitudinal Study of Women's Health (ALWSH) comparing women with (n=478) or without (n=8134) a self-reported diagnosis of PCOS. Main outcome measures were depression, anxiety and perceived stress measured using validated scales. The χ^2 and t tests were used to assess differences between groups. Univariable and multivariable regression were performed to determine factors contributing to each outcome.

Results. Women reporting PCOS, compared with women not reporting PCOS, reported higher prevalence of depression (27.3% ν . 18.8%), anxiety symptoms (50% ν . 39.2%) and greater score for perceived stress (1.01 ± 0.03 ν . 0.88 ± 0.01). After adjusting for body mass index, infertility and socio-demographic factors, women with PCOS were still more likely to be depressed, anxious and to have a higher level of perceived stress. There was a high-level mediation effect of stress between PCOS and both depression and anxiety.

Conclusion. Compared with women not reporting PCOS, women reporting PCOS have increased depression, anxiety and perceived stress. Stress may play a role in the association between PCOS, depression and anxiety. Further studies should consider assessment and management of stress in PCOS as it may be relevant for understanding the aetiology and treatment of psychological distress.

Introduction

Polycystic ovary syndrome (PCOS) is the most frequent endocrine disorder in women of reproductive age with a reported prevalence of 9–18% (March *et al.*, 2009). Its main clinical and diagnostic features include menstrual irregularity, biochemical and/or clinical hyperandrogenism and presence of polycystic ovaries on ultrasound (Aziz *et al.*, 2006). Women with PCOS can have severe metabolic and reproductive manifestations (Azziz *et al.*, 2004). PCOS is often also associated with excess body weight which can worsen the clinical presentation (Lim *et al.*, 2013). Compared with women without PCOS, women with PCOS show an increased prevalence of clinical depression (28–64% *v.* 7.1–8%), anxiety (34–57% *v.* 18%) (Deeks *et al.*, 2011) and other manifestations of psychological distress (Himelein and Thatcher, 2006; Kerchner *et al.*, 2009; Açmaz *et al.*, 2013; Rowlands *et al.*, 2016).

The aetiology of this observed increased prevalence of depression and anxiety in PCOS is still unclear. Possible explanatory factors that have been investigated as a source of distress include visible features such as excess weight, clinical hyperandrogenism (hirsutism, acne or androgenic alopecia) (Barry et al., 2011; Veltman-Verhulst et al., 2012), medical consequences such as infertility (Himelein and Thatcher, 2006; Tan et al., 2008; Deeks et al., 2010), concerns relating to diagnosis and fear regarding long-term health complications (Trent et al., 2003; Moran et al., 2010; Deeks et al., 2011). However, the contributing factors to depression and anxiety are currently unclear and results from studies are often conflicting. Only a few of the existing studies (Himelein and Thatcher, 2006; Benson et al., 2009b; Moran et al., 2010; Deeks et al., 2011) have adjusted for factors known to have an association with depression and anxiety in the general population such as body mass index (BMI) (Simon et al., 2006),

infertility (Cousineau and Domar, 2007) and socio-demographic factors such as ethnicity, household income and marital status (Moreno-Peral *et al.*, 2014). It is also well documented that a chronic illness is a stressful condition and chronic stress is associated with depression and anxiety in the general population (De Ridder *et al.*, 2008; Hammen *et al.*, 2009; Lahey, 2009). While women with PCOS reported a significantly increased physiological reaction to stress compared with controls (Benson *et al.*, 2009*a*; Farrell and Antoni, 2010), there is a paucity of studies investigating the relationship of stress with psychological health in PCOS (Guidi *et al.*, 2015). The prevalence of stress and its potential contribution to depression and anxiety in PCOS therefore warrants further investigation.

Furthermore, the majority of studies in PCOS have utilised clinic-based samples, convenience control groups, internet surveys (Benson et al., 2009a) or health service databases (Altinok et al., 2014; Hung et al., 2014). This limits the generalisability of their findings as referral bias may lead to findings of higher rates of overweight and obesity in such studies (Ezeh et al., 2013) and consequently higher rates of depression and anxiety as excess adiposity is related to worsened psychological health both in women with PCOS and in the general population (Simon et al., 2006; Álvarez-Blasco et al., 2010). The limited research in community-based studies in PCOS reports greater psychological distress in adolescents and young adults (Guidi et al., 2015; Rowlands et al., 2016) but none in adult women.

The aim of the present study was to assess depression, anxiety, perceived daily stress, psychotropic medication use and health-seeking behaviours with regards to managing psychological health in a large community-based, representative sample of Australian women, comparing women with a self-reported medical diagnosis of PCOS to those who did not report PCOS. We additionally aimed to evaluate the role of stress in contributing to the relation-ship between PCOS, depression and anxiety.

Methods

Study population and protocol

This study is based on data from the Australian Longitudinal Study on Women's Health (ALSWH), a longitudinal populationbased study of four age cohorts of Australian women. The study aims are to examine the physical and mental health of women across the life span assessing biological, psychological, social and lifestyle aspects as well as use and satisfaction with health care services. The ALSWH first collected mailed survey data from three age cohorts studied in 1996. Women were randomly selected from the national health insurance database Medicare (which includes all permanent residents of Australia) and invited to participate in the project. Women were recruited nationally with intentional oversampling from rural and remote areas (Lee et al., 2005). Further information about the methods used and sample characteristics has been reported elsewhere (Brown et al., 1999; Powers and Loxton, 2010) and are available on the ALSWH website (http:// www.alswh.org.au). The Human Research Ethics Committees of the University of Newcastle and the University of Queensland approved the study methods and informed written consent was obtained from each participant. The current study is based on data from the cohort born between 1973 and 1978 who first completed the survey at age 18-23 years. We analysed data from survey 4 (2006) and included women who responded to the question on PCOS (online Supplementary Fig. S1).

Demographic measures and participant characteristics

- (1) PCOS: Women were asked 'In the last three years, have you been diagnosed or treated for polycystic ovary syndrome?'. Women who responded 'yes' were classified as PCOS, whereas all other women who responded to the question were classified as not having PCOS.
- (2) BMI: Weight and height were self-reported. BMI (weight in kg/height in metres²) was calculated from self-reported height and weight. BMI was then classified into categories, with overweight and obesity defined by the World Health Organization criteria (BMI 25.0−29.9 kg/m² for overweight and ≥30 kg/m² for obese classification) (World Health Organization, 1999). Socio-demographic variables included age, education, occupation, marital status, number of children, ethnicity and personal income.
- (3) Fertility was assessed by the question 'Have you and your partner (current or previous) ever had problems with fertility that is, tried unsuccessfully for 12 months or more to get pregnant?' with the answers: (1) no never tried to get pregnant; (2) no, had no problem with fertility; (3) yes, but have not sought help/treatment and (4) yes, and have sought help/treatment available. Women who responded 'no never tried to get pregnant' or 'no, had no problem with fertility' were classified as not knowing/not experiencing infertility. Those who responded 'yes, but have not sought help/treatment' or 'yes, and have sought help/treatment' were classified as having experienced infertility.

Psychological outcomes

Both validated scales and more self-reported questions were used to investigate psychological features. Primary outcomes were depression and anxiety using standardised and validated tools.

- (4) Depression: Depression symptoms were assessed using the 10-item *Centre for Epidemiologic Studies Depression Scale* (CESD-10) with a score of ≥10 used as a categorical cut-off for clinically significant symptoms (Andresen *et al.*, 1994).
- (5) Anxiety: Anxiety symptoms were assessed using the anxiety subscale of the *Goldberg Depression and Anxiety Scale* (GADS) with a score of ≥5 used as a categorical cut-off for risk of clinical anxiety as previously described (Goldberg *et al.*, 1987, 1988).
- (6) Perceived stress: Perceived stress was analysed as a continuous variable assessed by the *Perceived Stress Questionnaire* (PPQ) (Bell and Lee, 2002, 2003) which is a scale that has been developed for and validated in the ALSWH. The total score ranges from 0 to 4 with a score >2 indicating moderate levels of stress and a score >3 or >4 indicating, respectively, a very and extremely stressful condition.

Secondary measures included self-reported information concerning medical diagnosis of mental illness, psychological symptoms and health-seeking behaviours relating to mental health management.

(7) Mental illness diagnosis: A self-reported medical diagnosis of depression, anxiety or other major mental illnesses was assessed with the question 'In the last three years have you been diagnosed or treated for'. Women could respond to these questions with either 'yes' or 'no' responses (yes/no).

(8) Psychological symptoms and seeking behaviours: Self-reported symptoms of depression, episodes of intense anxiety (e.g. panic attacks) or other mental problems and the proportion of women seeking help for those symptoms were investigated asking 'in the last 12 months have you had one of the following? (no, rarely, sometimes, often)'. Only women who responded 'often' were categorised as reporting symptoms. The question was then asked 'If yes, did you seek help for this problem? (yes/no)' and these data were also included in the analysis.

(9) Other health-seeking behaviours relating to mental health management: psychotropic medication use was investigated using the following two questions: 'During the past 4 weeks have you used medications that were: prescription medication for depression (e.g. Zoloft, Aropax, Lexapro, Cipramil, etc.), prescription medication for your nerves/anxiety (e.g. Valium, Serapax, Kalma, Ducene, etc.)'. Women could respond to these questions with either 'yes' or 'no' responses (yes/no). We also included the question 'Have you consulted a counsellor or other mental health worker for your own health in the last 12 months?' with '(yes/no)' responses in the analysis.

Statistical analysis

Data were analysed using Stata software version 11.0 (StataCorp, Texas, Lakeway, USA).

Categorical data were expressed as proportion and continuous data as mean ± standard deviation. Comparison between women with and without PCOS was performed by χ^2 for categorical variables and independent Student's t test for continuous variables. Binary logistic regression analyses were used to assess the relationship between depression or anxiety and PCOS, and linear regression models were used to assess the association between perceived stress and PCOS. Potential confounding variables in these relationships were assessed and incorporated in the model including socio-demographic factors, BMI (categorical variable) and infertility. The selection of variables was based on identifying all measured variables of known or suspected relevant effect on depression, anxiety and perceived stress, and/or exhibiting p < 10.10 on univariable analysis. In addition to examination as a dependent variable, perceived stress was also considered as a potential confounding variable in the association between PCOS, depression and anxiety. Since stress was also considered to be a mediator in the relationship between PCOS, depression and anxiety, binary mediation analysis was performed to partition the indirect effect of PCOS on both depression and anxiety. The significance level was set at a two-tailed 5%. Analysis was also weighted by area of residence to adjust for the deliberate oversampling in rural and remote areas.

Results

On 9145 responders to the survey, 8612 answered to the question of PCOS status and were included in the analysis (online Supplementary Fig. S1). Four hundred and seventy-eight were classified as reporting PCOS and 8134 as not reporting PCOS (abbreviated as with and without PCOS, respectively), with an estimated prevalence of self-reported PCOS of 5.8% (95% CI 5.3–6.3). Participant characteristics for the entire group and the PCOS and non-PCOS groups are reported in Table 1. The mean age was 30.6 ± 1.5 years and the mean BMI was 25.2 ± 5.7 kg/m². As previously reported (Teede *et al.*, 2013), women

with PCOS reported a greater BMI compared with women without PCOS (PCOS $28.1\pm7.2~{\rm kg/m^2}~\nu$. non-PCOS $25.1\pm5.6~{\rm kg/m^2}$) and a significantly higher proportion were in the overweight (PCOS $26.1\%~\nu$. non-PCOS 23.2%) and obese range (PCOS $33.2\%~\nu$. non-PCOS 16.2%) (p < 0.001). Women with PCOS were more likely to report experiencing infertility (PCOS $47.4\%~\nu$. non-PCOS 9.2%) (p < 0.001) (Table 1). Women with and without PCOS did not differ significantly in socio-demographic variables.

On unadjusted analysis, a significantly higher proportion of women with PCOS reported clinically significant depression (PCOS 27.3% v. non-PCOS 18.8%) and anxiety symptoms (PCOS 50.0% ν . non-PCOS 39.2%) (p < 0.001). Women with PCOS also had higher levels of perceived stress compared with women without PCOS (PCOS 1.06 ± 0.6 v. non-PCOS 0.88 ± 0.5) (p < 0.001) (Table 2). On univariable regression analysis, the odds of reporting depression symptoms or a score in the clinical risk range of depression was 1.61-fold higher in women with PCOS (95% CI 1.29–2.03, p < 0.001). The odds of reporting anxiety symptoms risk score was 1.46-fold higher in women with PCOS (95% CI 1.20–1.79, p < 0.001). The odds of reporting perceived stress was 0.16 in women with PCOS (95% CI 0.11-0.22, p < 0.001) (Table 2). Women with PCOS reported a significantly higher score than those without PCOS in all secondary psychological outcomes with the exception of seeking help for anxiety symptoms and other mental problems in the last 12 months. Women with PCOS were more likely to report psychotropic medication use both for depression (PCOS 12.6% v. non-PCOS 7.03%, p = 0.001) and anxiety (PCOS 5.4% ν . non-PCOS 2.8%, p = 0.001) (Table 1, online Supplementary Data).

In multivariable regression analysis, after adjusting for age, BMI, infertility, education, occupation, marital status, number of children, ethnicity and personal income, the effect size was slightly reduced for the outcomes of depression (OR 1.39, 95% CI 1.04–1.86, p=0.027), anxiety (OR 1.37, 95% CI 1.07–1.76, p=0.012) and perceived stress (coefficient β 0.15, 95% CI 0.09–0.23, p<0.001) (Tables 3–5). A second multivariable analysis was performed both for depression and anxiety, adding perceived stress to the other potential confounding variables described above. On these analyses, the association between PCOS with both depression and anxiety symptoms was no longer significant (Tables 4 and 5). On adjusted analyses, for a one-unit increase in stress, the odds of being clinically depressed and anxious increased by a factor of 9.93 and 11.36 (p<0.001), respectively.

On mediation analysis, we found that stress showed a high-level mediation effect between the relationship between PCOS and depression, with a proportion of total effect moderated = 0.71. The indirect effect of PCOS on depression was given by OR 1.18 (95% CI 0.92–1.51). A high-level mediation effect of stress between PCOS and anxiety was also found, with a proportion of total effect moderated = 0.71. The indirect effect of PCOS on anxiety was given by OR 1.18 (95% CI 0.95–1.47).

Discussion

The present study assessed for the first time depression and anxiety symptoms and perceived stress in adult women with PCOS in a large community-based cohort study. Women reporting PCOS reported an increased prevalence of depression and anxiety symptoms, perceived stress, self-reported medical diagnoses of depression, anxiety or other major mental illnesses and treatment for psychological conditions or mental illness. We also report here for the first time the confounding and the mediating effect of

Table 1. Characteristics for women with and without PCOS

Outcomes	Overall sample (n = 8612)	PCOS (n = 478)	non-PCOS (n = 8134)	p value
Age (years) (mean ± s.p.)	30.6 ± 1.5	30.5 ± 1.4	30.6 ± 1.5	0.127
BMI kg/m² (mean ± s.p.)	25.2 ± 5.7	28.1 ± 7.2	25.1 ± 5.6	<0.001
BMI WHO categories n (%)				<0.001
<25 kg/m ²	5139 (59.7)	183 (40.8)	4638 (60.5)	
25.0–29.9 kg/m²	2017 (23.4)	117 (26.1)	1780 (23.2)	
≽30 kg/m²	1456 (16.9)	149 (33.2)	1245 (16.2)	
Infertility n (%)				<0.001
Experienced infertility	1010 (11.1)	224 (47.4)	747 (9.2)	
Unknown/not experienced infertility	8105 (88.9)	249 (52.6)	7363 (90.8)	
Education n (%)				0.183
No formal qualification	90 (1.2)	9 (2.2)	78 (1.1)	
Year 12 or less	2211 (28.7)	107 (25.8)	1952 (24.8)	
Trade/certificate	292 (3.8)	14 (3.3)	259 (3.8)	
University/higher university degree	5120 (66.4)	284 (68.6)	4565 (66.6)	
Occupation n (%)				0.649
No job	2436 (27.2)	90 (19.2)	1565 (19.8)	
Professional, associate professional, manager	1740 (19.4)	211 (45.1)	3770 (47.3)	
Tradesperson, intermediate production or transport, labourer or related worker	4226 (47.2)	33 (7.1)	481 (6.1)	
Elementary, intermediate, advanced clerical	554 (6.2)	134 (28.6)	2157 (27.1)	
Marital status n (%)				0.562
Never married	2107 (23.1)	118 (24.8)	1850 (22.8)	
In couple (married, <i>de facto</i> opposite or same sex)	6587 (72.3)	339 (71.4)	5877 (72.5)	
Separated, divorced	401 (4.4)	18 (3.8)	366 (4.5)	
Widowed	11 (0.1)	0 (0.0)	11 (0.1)	
Number of children n (%)				0.356
No children	4688 (51.2)	258 (54.0)	4126 (50.7)	
1–2 children	3624 (39.6)	172 (36.0)	3260 (40.1)	
3–4 children	786 (8.6)	45 (9.4)	707 (8.7)	
5 or more children	47 (0.5)	3 (0.6)	41 (0.5)	
Ethnicity n (%)				0.596
Australian born	7572 (88.8)	414 (87.3)	7158 (88.9)	
Other English-speaking country born	542 (3.8)	24 (5.1)	352 (4.4)	
Europe	149 (1.1)	5 (1.1)	124 (1.5)	
Asia	381 (2.7)	24 (5.1)	322 (4.0)	
Other	134 (0.9)	7 (1.5)	99 (1.2)	
Average gross income per year Aus\$ n (%)				0.615
Not applicable	421 (4.7)	17 (3.7)	378 (4.8)	
0-15.999	2355 (26.6)	125 (26.9)	2117 (26.9)	
16.000-36.999	2225 (25.1)	117 (25.2)	1962 (24.9)	
37.000-77.999	3281 (37.1)	168 (36.2)	2910 (36.9)	
≽78.000	578 (6.5)	37 (7.9)	514 (6.5)	

Values are reported as mean (\pm s.b.) or number (%). Data were analysed by t test for continuous variables and χ^2 test for categorical variables. PCOS, polycystic ovary syndrome; BMI, body mass index; WHO, World Health Organization.

Table 2. Depression, anxiety and perceived stress in women with and without polycystic ovary syndrome

Outcome	PCOS	Non-PCOS	OR unadjusted (95% CI)	OR adjusted ^a	OR adjusted ^b
Depression CESD-10 n (%)	129 (27.3)	1505 (18.8)	1.61 (1.29–2.03)	1.39 (1.04–1.86)	1.04 (0.74–1.46)
Anxiety GADS n (%)	239 (50.0)	3175 (39.2)	1.46 (1.20-1.79)	1.37 (1.07–1.76)	1.02 (0.76–1.38)
Perceived stress PSQ mean ± s.p.	1.06 ± 0.61	0.88 ± 0.53	0.16 (0.11-0.22)	0.92 (0.22–3.79)	n/a

Values are reported as mean (±s.p.), number (%) or mean, 95% CI.

Data were analysed by t test for continuous variables and χ^2 test for categorical variables and survey-weighted univariable and multivariable logistic analysis.

stress in the association between both depression and anxiety and PCOS.

We report here an increased prevalence of clinical depression of 27.3% and 18.8% and anxiety of 50.0% and 39.2% in women with and without PCOS, respectively, compared with 7.1-8% and 18% in the general population (Deeks et al., 2011). In agreement with previous research (Barry et al., 2011; Dokras et al., 2011; Veltman-Verhulst et al., 2012), women with PCOS had increased risk of clinically significant depression and anxiety symptoms (1.39- and 1.37-fold respectively). These increased odds were lower than reported in a recent meta-analysis (4.03and 6.88-fold) (Dokras et al., 2011, 2012), which may be because the meta-analysis included clinical-based studies while our study used a community-based sample. Many factors in women may contribute to depression including hormonal and biological (e.g. infertility, child birth and premenstrual syndrome) and psychosocial (e.g. stress, socio-economic advantage and violence) factors (National Institute of Mental Health, 1995). We considered in our study BMI, infertility and socio-demographic variables as specific risk factors for depression for the general population (Simon et al., 2006). While some of these factors have been previously considered in investigating the relationship between PCOS and psychological variables (Himelein and Thatcher, 2006; Benson et al., 2009b; Moran et al., 2010; Deeks et al., 2011; Rowlands et al., 2016), to our knowledge, this is the first community-based study considering all of them together in one analysis.

In the present study, the association between PCOS, depression and anxiety was attenuated but maintained on adjustment for BMI, infertility and socio-demographic variables. This indicates that although the presence of overweight or obesity and infertility may worsen depression and anxiety as reported in the general population (Simon *et al.*, 2006), PCOS status *per se* is likely to have an independent effect on psychological function (Deeks *et al.*, 2010). This may be related to visible features, the frustration of having a chronic condition (Kozica *et al.*, 2013) or the perceived risk of future health complications (Moran *et al.*, 2010).

In agreement with prior results (Guidi et al., 2015), being stressed was more prevalent in PCOS even after consideration of BMI, infertility and socio-demographic variables. The questionnaire used here assesses perceived sources of stress in several domains of daily life (Bell and Lee, 2002, 2003) rather than being designed for the assessment of health-related stress. However, it also assesses perceived stress in life domains potentially impacted

by PCOS such as health, motherhood or social relationships (Farkas et al., 2014). Future work should investigate if women with PCOS display higher stress related to their condition rather than other daily life domains or if stress precedes PCOS. We also report here for the first time the role of stress both as confounder and mediator variable between the relationship between PCOS and psychological outcomes. The significant association between PCOS and depression and anxiety was lost on adjustment for stress, and stress showed a strong association both with depression and anxiety. This suggests an independent confounding relationship of stress both with PCOS status and with depression or anxiety and that higher depression and anxiety in PCOS may be related to higher stress levels in PCOS. Mediation analysis gave further clarification as stress mediated for a large proportion of the relationship between PCOS and both depression and anxiety. This could suggest that stress can have a relevant direct effect in depression and anxiety symptoms in women with PCOS, rather than other factors considered up to now. This is consistent with prior research on the relationship between stress, chronic illness and psychological morbidity in the general population (Hammen et al., 2009; Lahey, 2009). This may explain why previous studies have not found any direct causal relationship between PCOS and depression or anxiety.

As endocrine systems may be more vulnerable to the physiological effects of stress due to the pathophysiological features of PCOS such as hypothalamic-pituitary-adrenal axis (HPA) and sympathetic nervous system (SNS) hyperactivity or low-grade immune system inflammation (Benson *et al.*, 2008), even low levels of perceived stress may have a clinically significant impact in PCOS (Farrell and Antoni, 2010; Barry *et al.*, 2011). In light of this, the evaluation of coping and adjustment strategies in PCOS is relevant as key factors relating to psychological outcomes (Benson *et al.*, 2010) and preliminary trials of stress management interventions have reported encouraging results on the amelioration of psychological outcomes in PCOS (Raja-Khan *et al.*, 2015; Stefanaki *et al.*, 2015).

The findings of the present study therefore confirm the importance of consideration of assessment and management of stress, even at non-clinical levels, in PCOS in addition to depression and anxiety. Furthermore, the independent association of BMI and infertility with both depression and anxiety was also lost on inclusion of stress into the multivariable models. Stress may therefore have a stronger impact on psychological distress than other factors previously highlighted as risk factors for

PCOS, polycystic ovary syndrome; CESD-10, Centre for Epidemiologic Studies Depression Scale; GADS, Goldberg Anxiety Depression Scale; PSQ, Perceived Stress Questionnaire.

^aAdjusted for BMI, infertility and socio-demographic factors (age, education, occupation, marital status, number of children, ethnicity, personal income).

^bAdjusted for perceived stress, BMI, infertility, socio-demographic factors (age, education, occupation, marital status, number of children, ethnicity, personal income).

Table 3. Univariable and multivariable regression analysis – depression

	OR (95% CI) unadjusted	OR (95% CI) adjusted ^a	OR (95% CI) adjusted ^b
PCOS	1.61 (1.29–2.03)	1.38 (1.03–1.86)	1.04 (0.74–1.46)
Stress	10.32 (9.01–11.83)	-	9.9 (9.93–11.74)
Age (years)	0.99 (0.95–1.03)	1.00 (0.95–1.05)	1.01 (0.96–1.07)
BMI WHO categories			
<25.0 kg/m ²		1	
25.0–29.9 kg/m²	1.25 (1.08–1.45)	1.24 (1.05–1.47)	1.17 (0.96–1.42)
≥30 kg/m²	1.93 (1.66–2.25)	1.61 (1.35–1.92)	1.24 (1.01–1.55)
Experienced infertility	1.26 (1.06–1.50)	1.27 (1.01–1.60)	1.21 (0.93–1.58)
Education			
No formal qualification		1	
Year 12 or less	0.50 (0.31-0.81)	0.64 (0.34–1.22)	0.47 (0.22-0.99)
Trade/certificate	0.34 (0.19-0.62)	0.47 (0.22-0.99)	0.31 (0.13-0.71)
University/higher university degree	0.36 (0.22–0.57)	0.52 (0.27-0.99)	0.36 (0.17-0.75)
Occupation			
No job		1	
Professional, associate professional, manager	0.70 (0.60-0.82)	0.77 (0.60-0.99)	0.67 (0.51-0.90)
Tradesperson, intermediate production or transport, labourer or related worker	1.28 (0.99–1.66)	1.15 (0.83–1.59)	1.03 (0.72–1.48)
Elementary, intermediate, advanced clerical	1.11 (0.93–1.30)	0.99 (0.79–1.25)	0.84 (0.64–1.10)
Marital status			
Never married		1	
In couple (married, de facto opposite or same sex)	0.60 (0.49-0.64)	0.54 (0.45-0.65)	0.62 (0.51-0.77)
Separated, divorced	1.90 (1.48-2.43)	1.68 (1.23-2.30)	1.21 (0.83-1.78)
Widowed	2.94 (0.79–10.98)	8.41 (1.60-44.39)	6.55 (1.49–28.72)
Number of children			
No children			
1–2 children	0.91 (0.81–1.03)	0.82 (0.67-0.99)	0.72 (0.57–0.89)
3-4 children	1.20 (0.98–1.47)	0.84 (0.63-1.12)	0.66 (0.48-0.90)
5 or more children	2.52 (1.30–4.90)	2.14 (0.96–4.80)	2.30 (0.92–5.81)
Ethnicity			
Australian born		1	
Other English-speaking country born	1.21 (0.93–1.58)	1.03 (0.74–1.44)	1.06 (0.73–1.53) p =
Europe	0.87 (0.52–1.45)	0.98 (0.55–1.74) p =	1.40 (0.71–2.78) p =
Asia	0.90 (0.66–1.22)	0.90 (0.62–1.32)	1.02 (0.66–1.57)
Other	0.53 (0.28–0.96)	0.77 (0.38–1.55)	0.88 (0.36-2.14)
Average gross income per year Aus\$ n (%)			
Don't know/don't want to tell		1	
0-15.999	0.78 (0.60–1.03)	0.97 (0.68–1.40)	0.92 (0.59–1.43)
16.000–36.999	0.91 (0.69–1.21)	0.94 (0.66–1.35)	0.78 (0.50-1.22)
37.000–77.999	0.67 (0.51-0.88)	0.86 (0.59–1.26)	0.81 (0.51–1.27)
≽78.000	0.48 (0.33-0.69)	0.72 (0.44-1.18)	0.75 (0.42-1.33)

Data were analysed by survey-weighted univariable and multivariable logistic analysis.

PCOS, polycystic ovary syndrome; BMI, body mass index; WHO, World Health Organization.

^aAdjusted for BMI, infertility and socio-demographic factors (age, ethnicity, education, marital status, number of children, personal income).

^bAdjusted for perceived stress, BMI, infertility, socio-demographic factors (age, ethnicity, education, marital status, number of children, personal income).

Table 4. Univariable and multivariable regression analysis – anxiety

	OR (95% CI) unadjusted	OR (95% CI) adjusted ^a	OR (95% CI) adjusted ^b
PCOS	1.46 (1.20–1.79)	1.37 (1.07–1.76)	1.02 (0.76-1.37)
Stress	10.32 (9.20-11.57)	-	11.36 (9.64–13.40
Age	0.98 (0.95-1.01)	0.97 (0.94–1.01)	0.97 (0.93-1.02)
BMI WHO categories n (%)			
<25 kg/m ²		1	
25-0-29.9 kg/m ²	1.01 (0.90-1.13)	0.98 (0.85-1.12)	0.88 (0.75-1.03)
≥30 kg/m²	1.60 (0.90-1.13)	1.34 (1.15–1.56)	1.03 (0.86-1.23)
Infertility	1.28 (1.11-1.49)	1.13 (0.94–1.36)	1.07 (0.88-1.33)
Education			
No formal education		1	
Secondary school (up to 10–12 years)	0.62 (0.38-1.01)	0.57 (0.31–1.06)	0.44 (0.24-0.82)
Certificate diploma	0.62 (0.36–1.07)	0.57 (0.29–1.13)	0.41 (0.21-0.82)
Tertiary school (trader qualification, degree, higher degree)	0.47 (0.29–0.76)	0.45 (0.24-0.84)	0.32 (0.17-0.60)
Occupation			
No job		1	
Professional, associate professional, manager	1.45 (1.19–1.77)	1.04 (0.85-1.28)	0.99 (0.78-1.24)
Tradesperson, intermediate production or transport, labourer or related worker	1.39 (1.25–1.56)	1.17 (0.89–1.52)	1.12 (0.81–1.53)
Elementary, intermediate, advanced clerical	1.19 (1.05-1.34)	1.23 (1.02-1.48)	1.16 (0.94–1.42)
Marital status			
Never married		1	
In couple (married, de facto opposite or same sex)	0.76 (0.68-0.85)	0.76 (0.66-0.88)	0.92 (0.77-1.10)
Separated, divorced	1.65 (1.30-2.09)	1.46 (1.10-1.96)	0.98 (0.70-1.10)
Widowed	1.54 (0.41-5.79)	5.99 (1.02-35.16)	3.76 (1.02–13.79
Number of children			
No children		1	
1–2 children	0.91 (0.81-1.03)	0.94 (0.81–1.10)	0.90 (0.76-1.06)
3–4 children	1.20 (0.98-1.47)	0.99 (0.78–1.24)	0.84 (0.65–1.09)
5 or more children	2.52 (1.30-4.90)	1.37 (0.62-3.03)	1.41 (0.57-3.45)
Ethnicity n (%)			
Australian born		1	
Other English-speaking country born	0.95 (0.76–1.19) p = 0.7	0.87 (0.66–1.15)	0.86 (0.60-1.23)
Europe	0.74 (0.50-1.10)	0.83 (0.52-1.31)	1.09 (0.60-1.23)
Asia	1.16 (0.93–1.47)	1.14 (0.86–1.51)	1.33 (0.96–1.83)
Other	0.62 (0.40-0.96)	0.62 (0.36–1.05)	0.65 (0.37-1.15)
Average gross income per year Aus\$ n (%)			
Don't know/don't want to tell		1	
0-15.999	0.93 (0.74–1.18)	0.91 (0.68-1.22)	0.79 (0.57-1.11)
16.000-36.999	1.08 (0.86-1.37)	0.92 (0.68-1.22)	0.72 (0.51-1.01)
37.000-77.999	0.84 (0.67–1.06)	0.82 (0.61–1.11)	0.71 (0.50-1.00)
≽78.000	0.75 (0.57–0.99)	0.81 (0.56–1.19)	0.79 (0.51-1.21)

 $\label{lem:decomposition} \mbox{Data were analysed by survey-weighted univariable and multivariable logistic analysis.}$

PCOS, polycystic ovary syndrome; BMI, body mass index; WHO, World Health Organization.

^aAdjusted for BMI, infertility and socio-demographic factors (age, ethnicity, education, marital status, number of children, personal income).

^bAdjusted for perceived stress, BMI, infertility, socio-demographic factors (age, ethnicity, education, marital status, number of children, personal income).

Table 5. Univariable and multivariable regression analysis – stress

	eta-coefficient (95% CI) unadjusted	eta-coefficient (95% CI) adjusted
PCOS	0.15 (0.11–0.22)	0.16 (0.09-0.23)
Age (years)	-0.01 (-0.01 to 0.01)	-0.01 (-0.01 to 0.01)
BMI WHO categories		
<25 kg/m ²	1	
25.0–29.9 kg/m ²	0.04 (0.02-0.07)	0.04 (0.01-0.07)
≽30 kg/m²	0.17 (0.13-0.20)	0.14 (0.10-0.18)
Infertility	0.05 (0.01-09)	0.03 (-0.02 to 0.07)
Education		
No formal education	1	
Secondary school (up to 10–12 years)	0.02 (-0.11 to 0.15)	0.07 (-0.07 to 0.21)
Certificate diploma	0.02 (-0.12 to 0.17)	0.10 (-0.06 to 0.26)
Tertiary school (trader qualification, degree, higher degree)	0.01 (-0.12 to 0.13)	0.08 (-0.06 to 0.22)
Occupation		
No job	1	
Professional, associate professional, manager	-0.003 (-0.04 to 0.14)	0.03 (-0.02 to 0.08)
Tradesperson, intermediate production or transport, labourer or related worker	0.08 (0.01-0.14)	0.04 (-0.02 to 0.11)
Elementary, intermediate, advanced clerical	0.07 (0.03-0.10)	0.05 (0.01-0.10)
Marital status		
Never married	1	
In couple (married, de facto opposite or same sex)	−0.11 (−0.14 to −0.08)	-0.11 (-0.15 to 0.07)
Separated, divorced	0.25 (0.18-0.32)	0.21 (0.13-0.29)
Widowed	0.27 (-0.06 to 0.59)	0.30 (0.01-0.61)
Number of children		
No children	1	
1–2 children	0.01 (-0.02 to 0.03)	0.02 (-0.02 to 0.06)
3–4 children	0.06 (0.01-0.10)	0.08 (0.01-0.14)
5 or more children	0.06 (-0.16 to 0.27)	0.06 (-0.17 to 0.29)
Ethnicity		
Australian born	1	
Other English-speaking country born	-0.01 (-0.07 to 0.05)	-0.01 (-0.09 to 0.07)
Europe	-0.09 (-0.18 to -0.01)	-0.14 (-0.23 to -0.05)
Asia	-0.01 (-0.07 to 0.5)	-0.03 (-0.11 to 0.36)
Other	-0.08 (0.17-0.91)	-0.05 (-0.15 to 0.05)
Average gross income per year Aus\$ n (%)		
Don't know/don't want to tell	1	
0-15.999	0.01 (-0.06 to 0.07)	0.36 (-0.04 to 0.11)
16.000-36.999	0.10 (0.04–0.17)	0.08 (0.01–0.16)
37.000-77.999	0.01 (-0.05 to 0.07)	0.03 (-0.05 to 0.10)
≽78.000	-0.06 (-0.13 to 0.01)	-0.02 (-0.11 to 0.07)

Data were analysed by survey-weighted univariable and multivariable regression analysis.

PCOS, polycystic ovary syndrome; BMI, body mass index; WHO, World Health Organization.

^aAdjusted for BMI, infertility and socio-demographic factors (age, ethnicity, education, marital status, number of children, personal income).

depression and anxiety both in PCOS and in the general population.

We report here a more comprehensive assessment of psychological distress in women with and without PCOS than previously reported. A significantly higher proportion of women with PCOS reported having been diagnosed or treated for depression, anxiety or other mental illness, used psychotropic medications and suffered from or sought help for psychological distress symptoms or consulted a mental health professional. This is in keeping with the higher prevalence of depression and anxiety in PCOS. However, the self-reported prevalence of symptoms or episodes of depression or anxiety and the percentage of women engaged in psychological health-seeking behaviours was also lower than the prevalence of depression and anxiety symptoms using validated scales. This suggests that further research is warranted investigating if women with PCOS are aware of treatment options and strategies to encourage them to seek help for psychological distress. While there are numerous studies assessing psychological distress in women with PCOS, there is relatively limited literature either examining health-seeking behaviours relating to mental health management (Benson et al., 2009b; Moran et al., 2010; Altinok et al., 2014) or psychological treatments (Rofey et al., 2008; Raja-Khan et al., 2015) either in clinical or community samples. As far as we know that is only the second study (Altinok et al., 2014) investigating the use of psychotropic medication among women with PCOS. In accordance with these previous results, we found that women with PCOS were more likely to use psychotropic medication than women without PCOS. This is an important aspect of health-seeking behaviours that need to be further explored.

There is therefore a need to report community-based data on current uptake of management strategies for psychological health. This may encourage clinicians to discuss management options with patients for psychological features.

Strengths of this study include the use of a community-based cohort which reduces the bias of clinical-based studies in overestimating overweight and obesity (Ezeh et al., 2013), reproductive (Khan et al., 1996) and potentially psychological issues. Further strengths include the consideration of a broad range of potential confounding variables, the use of valid and reliable psychological measures as primary outcomes, the assessment of multiple less commonly investigated psychological outcomes and the examination of the effect of stress in psychological distress in women with PCOS. This is a cross-sectional analysis and neither causality nor a longitudinal evaluation between PCOS and psychological distress can be established. The major limitations of this study are the use of self-reported information such as PCOS status, infertility, BMI and psychological outcomes. However, this is reasonable in large epidemiologic studies due to feasibility and economy. Furthermore, validation studies have previously reported the validity of self-report measures for anthropometric measures as well as medical and psychological records in the general population or primary care setting (Von Korff et al., 1996; Haapanen et al., 1997; Spencer et al., 2002). Another limitation is the lack of information about treatments such as antiandrogen therapy and contraception which may have potential effect on mood (Davis and Tran, 2001; Oinonen and Mazmanian, 2002). The women in this study were also all aged between 28 and 33 years, and these results may not be applicable to women in other age groups. This is the first study assessing depression, anxiety symptoms and perceived stress in adult women with and without PCOS in a large, unselected community cohort. This

study confirms findings of worsened psychological distress and highlights the role of even non-clinical stress levels in potentially playing a role in the association among PCOS, depression and anxiety. It emphasises the need for a multidisciplinary approach to this increasing and debilitating condition. Future work should consider more specific assessments of health-related stress in women with PCOS and the potential role for stress management in the treatment of psychological distress in PCOS.

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