
1 **A Systematic Review and Meta-analysis of Polycystic Ovary Syndrome and**
2 **Mental Health among Black Asian Minority Ethnic populations**

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4 Authors

5 Gayathri DELANEROLLE, ^{1*}, Salma AYIS^{2*}, Vanya BARZILOVA^{3**}, Peter
6 PHIRI,^{5,7***}, Yutian ZENG^{9**}, Sandali RANAWEERA,^{6**}, Ashish SHETTY^{3,4**}, Nyla
7 HAQUE¹², Debasish KAR¹, Kingshuk MAJUMDER⁸, Shanaya RATHOD⁵, Vanessa
8 RAYMONT,¹², Jian Qing SHI^{9,10***}, Dharani K. HAPANGAMA^{3,7***}

9
10 *Affiliations*

11 ¹University of Oxford, Nuffield Department of Primary Health Care Sciences

12 ²Kings College London

13 ³University of Liverpool

14 ⁴University College London Hospitals NHS Foundation Trust

15 ⁵University College London

16 ⁶Southern Health NHS Foundation Trust

17 ⁷Liverpool Women's NHS Foundation Trust

18 ⁸University of Southampton, School of Primary Care, Population Sciences and
19 Medical Education, Faculty of Medicine

20 ⁹University of Manchester NHS Foundation Trust

21 ¹⁰Southern University of Science and Technology

22 ¹¹Alan Turing Institute

23 ¹²Department of Psychiatry, University of Oxford

24

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52 **Correspondence to:** dharani@liv.ac.uk

53 Professor Dharani K. Hapangama

54 Department of Women's and Children's Health,

55 Institute of Life Course and Medical Sciences,

56 University of Liverpool

57 Liverpool Women's Hospital NHS Trust

58 Crown Street, Liverpool L8 7SS, United Kingdom

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118 **Abstract**

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120 **Background:** Polycystic Ovary Syndrome (PCOS) is a chronic and a common
121 gynaecological condition impacting women of reproductive age. Women with PCOS

122 have hormonal, ovulatory and metabolic dysfunction resulting in multiple symptoms.
123 The correlation between hormonal disbalance and the impact on women's mental
124 health (MH) has been researched for decades. However, the prevalence among
125 different ethnicities has not been fully evaluated.

126

127 **Methods:** A systematic methodology was developed, and a protocol was published
128 in PROSPERO (CRD42020210863) and a systematic review of publications
129 between 1st January 1990- 30th January 2021 was conducted. Multiple electronic
130 databases were explored using keywords and MeSH terms. The finalised dataset
131 was analysed using statistical methods such as random-effect models, subgroup
132 analysis and sensitivity analysis.

133

134 **Findings:** We included 30 studies reporting on 3,944 PCOS women. Majority of
135 studies addressed depression anxiety, and common mental health. Studies had fair
136 to poor methodological quality and includes observational studies and Randomised
137 Clinical Trials (RCTs). Overall, 17% (95% CI: 7% to 29%) of women with PCOS have
138 clinical diagnosis of major or severe depression; 33% (95% CI: 26% to 40%) have
139 elevated depressive symptoms or a clinical diagnosis of depression; 41% (95% CI:
140 28% to 54%) report anxiety symptoms, and 31% (95% CI: 15% to 51%) have a form
141 a common mental health or are taking psychiatric medication for anxiety and / or
142 depression. The use of various tools to assess mental health symptoms was among
143 the reasons for the substantial heterogeneity across studies.

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145

146 **Interpretation:**

147 PCOS is associated with an increased risk of mental health disorders including
148 depressive s, anxiety, and other mental health disorders. While BAME populations
149 account for about 20% of most of the samples studied, stratification by ethnicity was
150 rarely attempted which made it difficult to elucidate the MH impact of PCOS on
151 different communities.

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153

154 **Keywords: Polycystic Ovary Syndrome, PCOS, BAME, Mental Health, Women's**

155 **Health and Wellbeing**

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177 **Introduction**

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179 Polycystic ovary syndrome (PCOS) is the most common endocrine disorder in
180 women of reproductive age with significant reproductive, metabolic, and
181 psychological implications. An estimated 8-13% of women of reproductive age are

182 affected,¹⁻⁵ with a widely quoted similar prevalence between different ethnicities.⁶
183 The Rotterdam criteria,^{5,7} which incorporates a combination of signs and symptoms
184 of androgen excess, ovarian dysfunction and polycystic ovarian morphology on
185 ultrasound is the globally accepted diagnostic criteria for PCOS. The phenotype
186 between different ethnicities, can vary markedly^{5,8,9} with PCOS sufferers from
187 Hispanic, East Asian, South Asian and Middle Eastern backgrounds displaying
188 higher prevalence of symptoms associated with hyperandrogenism^{8,9} than their
189 White counterparts. The precise aetiology of the syndrome is unclear, although
190 evidence suggests a genetic predisposition, and a role of environmental variables
191 including diet and lifestyle factors.^{10,11} The pathophysiology involves a dysfunctional
192 hypothalamic-pituitary-ovarian axis, with inappropriate gonadotropin-releasing
193 hormone (GnRH) pulsatility and increased pituitary secretion of luteinizing hormone
194 (LH).¹² Those affected with PCOS present with a wide spectrum of clinical symptoms
195 and features, which can be reproductive (oligomenorrhoea, hirsutism, subfertility),
196 metabolic^{13,14} (insulin resistance, type 2 diabetes, metabolic syndrome), and
197 psychological^{15,16} (depression, anxiety and low quality of life).
198
199 PCOS, due to its diversity in clinical presentation, remains undiagnosed in up to 70%
200 of women,³ this may lead to various clinical sequelae that impact and complicate
201 women's long-term health. For example women with PCOS have an increased risk
202 of consequential complications such as type 2 diabetes mellitus, cardiovascular
203 disease, endometrial cancer, pregnancy complications and psychological
204 conditions.¹⁷⁻²⁰
205
206 One area lacking in PCOS research are the MH correlates of the syndrome. The
207 limited evidence available suggests that women with PCOS have higher rates of

208 depression, anxiety, binge eating disorders, bipolar disorder and other psychological
209 symptoms,²¹⁻²³ most likely due to the androgenic, reproductive, and metabolic
210 disorders or due to the associated symptoms of the disorder.^{24,25} Current treatment
211 for PCOS focuses on physical symptom control and reducing complication rates.²⁶ At
212 a global level, currently, there is limited recognition of mental health (MH) symptoms
213 in women with PCOS.⁵ However, background MH sequelae may significantly impact
214 treatment adherence and patient quality of life.⁵

215

216 MH disorders account for 14% of global burden of disease.²⁷ Yet significant
217 disparities persist in providing access to MH services for Black, Asian and Minority
218 Ethnic (BAME) populations.²⁸ This is further exacerbated by the underreporting of
219 MH symptoms in BAME individuals who are less likely to contact their general
220 practitioner regarding their MH compared to their White British counterparts.²⁸ This
221 can be explained by a combination of personal and environmental factors, including
222 inability to recognise MH symptoms, cultural identity and stigma, language barrier
223 and poor communication with healthcare professionals.²⁸ Therefore, the prevalence
224 of MH disturbances in BAME women with PCOS remains unclear. The MH symptom
225 prevalence rates may differ significantly across races and ethnicities for women with
226 PCOS in the form of either symptoms and/or psychiatric disorders. Understanding
227 the multifactorial and heterogenic nature of PCOS presentation in BAME women is
228 crucial to improve clinical and patient reported outcomes.

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230 An assessment of the existing evidence is vital to generate comprehensive
231 knowledge and clinical practice improvements that are suitable for BAME

232 populations. Therefore, a systematic review was designed and developed within this
233 study.

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241 **Methods**

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243 A systematic methodology was developed and peer reviewed prior to publication in
244 PROSPERO in accordance with Preferred Reporting Items for Systematic Reviews
245 and Meta-Analyses (PRISMA).

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247 *Research question*

248 To assess the prevalence of the disease sequelae between PCOS and mental
249 health symptoms and disorders among BAME populations.

250

251 *Search Strategy*

252 The search strategy involved multiple databases of EMBASE, PubMed, PsycINFO,
253 Science direct and Web of Science from inception to the 30th of May 2021. The
254 searches were conducted through references initially followed by the review of their
255 abstracts and tracking of citations.

256

257 *Eligibility criteria*

258 All peer review publications in English were included to this study. Key words and
259 MeSH terms included *depression, polycystic ovary syndrome, PCOS, bipolar,*
260 *dysthymia, major depressive disorder, anxiety, psychosis, somatic symptoms,*

261 *psychotic disorders, trauma, stress, post traumatic stress (PTSD), obsessive-*
262 *compulsive spectrum disorders, ethnic minorities, BAME, BME and eating disorders.*
263 Studies that included a psychiatric diagnosis or MH symptomatologies based on a
264 clinically structured diagnostic interview and/or clinically accepted screening tools
265 were eligible for this study. Meeting abstracts were excluded if there was no full
266 paper available.

267

268 *Data extraction and analysis*

269 Predefined clinical variables were used as part of the data extraction of rates of
270 depression, anxiety, stress, schizophrenia, psychosis and PTSD within the search
271 strategy. An evidence synthesis methods protocol was developed and has been
272 shown as supplementary document 1. The data extraction process was completed
273 as per the PRISMA guidelines whilst, the refinement protocol has been shown in the
274 supplementary document 1. The extracted data was reviewed using Endnote and
275 Microsoft Excel by 3 reviewers prior to the statistical analysis. All studies collated
276 were categorised based on MH symptoms versus psychiatric disorders and, where
277 possible synthesised based on prevalence and 95% confidence intervals.
278 Prevalence tables were developed to indicate any subgroup categories including
279 geographical location, race, heterogeneity, age, obesity scores and ethnicity.
280 Systematically included studies that did not meet the meta-analysis criteria due to
281 insufficient statistical data or poor quality were narratively synthesised and analysed
282 based on the disease, family, clinician and patient perspective. The narration would
283 include the reporting of common themes including potential barriers to identify and
284 report any themes or sub-themes that may be present in clinical guidelines.

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289 *Outcomes*

290 The primary outcome was to assess the odds ratios and p-values associated with a
291 possible mental health symptomatology and/or psychiatric disorders among PCOS
292 BAME populations.

293

294 *Statistical analysis plan*

295 Depression was measured by a range of scales and several tools including the
296 Diagnostic and Statistical Manual of Mental Disorders (DSM)²⁹; the Patient Health
297 Questionnaire (PHQ)³⁰; the Hospital Anxiety and Depression (HAD)³¹; Beck
298 Depression Inventory (BDI)³²; Montgomery Åsberg Depression Rating Scale
299 (MADRS-S)³³; Center for Epidemiologic Studies Depression (CES-D)³⁴; Reynolds
300 Adolescent Depression Scale, 2nd Edition (RADS-2)³⁵; Quick Inventory Depressive
301 Symptomatology (QIDS)³⁶; Self-Reported Questionnaire Scale-20 (SRQ-20)³⁷; the
302 Generalized Anxiety Disorder (GAD)³⁸; the International Neuropsychiatric Interview
303 (MINI),³⁹ use of antidepressants or use of anti-anxiety medication; a diagnosis by a
304 MH professional, and self-reported history of depression.

305

306 The types of mental illness considered by most studies were anxiety, depression,
307 both anxiety and depression, and generalised mental wellbeing. The prevalence of
308 these was synthesised using a meta-analysis.

309

310 For the purpose of this study, depression was defined as a major depression
311 diagnosis or elevated depressive symptoms, as reported within the systematically
312 included studies. The diagnosis of major depressive disorder (MDD) was based on
313 clinical review which included suicidal ideations, or provided a definition of MDD
314 based on thresholds of tools used, for example, BDI ≥ 17 , CES-D ≥ 24 .^{40,41} Elevated
315 depression included clinical depression, antidepressants use, and / or a threshold for
316 mild to moderate depression based on the scale used for assessment, for example
317 BDI ≥ 11 , HAD-D ≥ 8 , and CES-D ≥ 16 . For anxiety, a combinational approach was
318 used where the use of the term anxiety or use of anxiety medications were pooled
319 together, due to overlapping definitions, and lack of clear cuts for severity levels.
320 Generalised mental wellbeing on the other hand was defined, as having both
321 depression and anxiety, use of medication for psychiatric disorders as reported
322 within the studies, self-reported MH symptoms, or reported under the term “common
323 mental health” (CMH) by a study authors based on scales such as SF-36.⁴²⁻⁴⁴

324

325 A Random-effects models were used to calculate overall summary estimates of
326 anxiety, depression, and CMH in PCOS patients.⁴⁵ The routine “metaprop” of the
327 software STATA’s (V.16) which provides pooled summary of proportions and 95%
328 confidence intervals based on Newcombe exact binomial estimation was used.^{46,47}

329 The routine also estimates the I-squared statistics which describes the percentage of

330 variation across studies that is due to heterogeneity rather than chance. (Higgins et
331 al. 2003)⁴⁸

332

333 All studies systematically included demonstrate a range of study designs including of
334 observational and randomised clinical trials (RCTs). Where comparisons were made
335 between PCOS and non-CPOS groups in a study, only the PCOS groups were
336 included. At situations where PCOS population were stratified by certain criterion, for
337 example body mass Index (BMI) or age, the strata were combined where the sample
338 size was smaller than 30 for each group, and an overall prevalence was calculated.
339 For larger groups, each was treated a separate group. Sensitivity analyses in
340 addition, were used to highlight differences across strata where a criterion proven to
341 have an impact on heterogeneity.

342

343 Publication bias was assessed using funnel plots and Egger test, for symmetry and
344 small study effects where the number of studies exceeded the recommended 10.^{49,50}

345

346 *Risk of Bias (Quality assessment)*

347 The Newcastle-Ottawa-Scale (NOS) was used to assess the risk of bias (RoB) for all
348 systematically included studies as demonstrated within the RoB table.

349

350 *Sensitivity analysis*

351 Sensitivity analysis was used to explore causes of heterogeneity. BMI, and tool of
352 assessment of assessments were used for stratification.

353

354

355

356 **Results**

357

358 Figure 1 displays the estimated prevalence and 95% confidence intervals (CI) for the
359 mental health conditions classified as: (a) Major depressive disorder, (b) Elevated
360 existing depression, (c) Anxiety and (d) Common mental health.

361

362 Overall, major depressive disorder affected 17% of PCOS patients, and vary
363 between 0% to 67%. All studies except two have estimated a prevalence \geq 10%. The
364 highest prevalence was reported in a Swedish study, followed by two Middle
365 Eastern (ME) studies, with a prevalence of 67%, 29%, and 22% for the three studies
366 respectively.⁵¹⁻⁵³ Substantial heterogeneity, I^2 of 93.98% was reported between
367 regions, also considerably high within USA studies, I^2 of 71.33% but the two ME
368 studies were homogeneous, Figure 1 (a).

369

370 Depression was summarised from 24 studies, including 3157 women with PCOS.
371 The estimated average was 33% (95% CI: 26% to 40%). Where the estimated
372 prevalence was stratified by region, fairly similar estimates on average, were noted
373 across regions, with India being the only exception where the the highest prevalence
374 of 64% (95% CI: 55% to 73%) was reported. Substantial heterogeneity observed
375 within regions and between regions. The overall heterogeneity was 93.24%.

376

377 For anxiety, Figure 1 (C), the highest prevalence of 75% was seen in a USA study,
378 with 78% white and 22% non-white, mean age 33 years (SD: 7.5), followed by a
379 prevalence of 74% in a European study with 80% white, 18% Asian, age 33 years
380 (7.4), while the two studies didn't report BMI.^{54,55} Middle East, and Australia/ NZ

381 follow with an estimated averages of 42% and 38% respectively, and the overall
 382 estimated prevalence was 41% (95% CI: 28% to 54%).

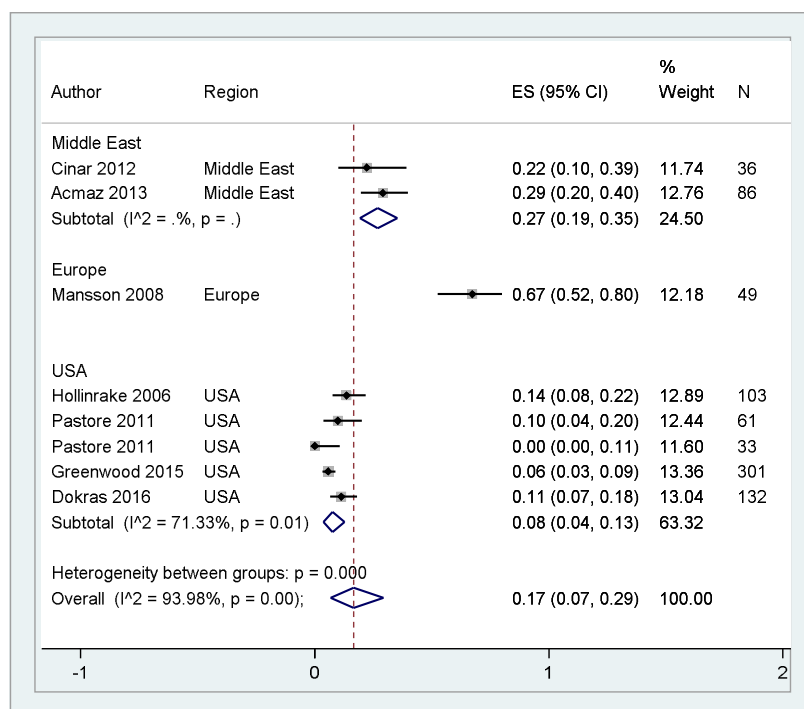
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384 Figure 1 (d), common mental health overall prevalence was 31% (95% CI: 15% to
 385 51%). Substantial heterogeneity shown by USA studies, only one study from ME,
 386 and one from Brazil were included. A substantial lack of consistency across studies
 387 was reflected by a heterogeneity I^2 of 97.29%.

388

389 Figure 1. The Prevalence of Mental Health Indicators in Women with PCOS

390 (a) Major depression by region

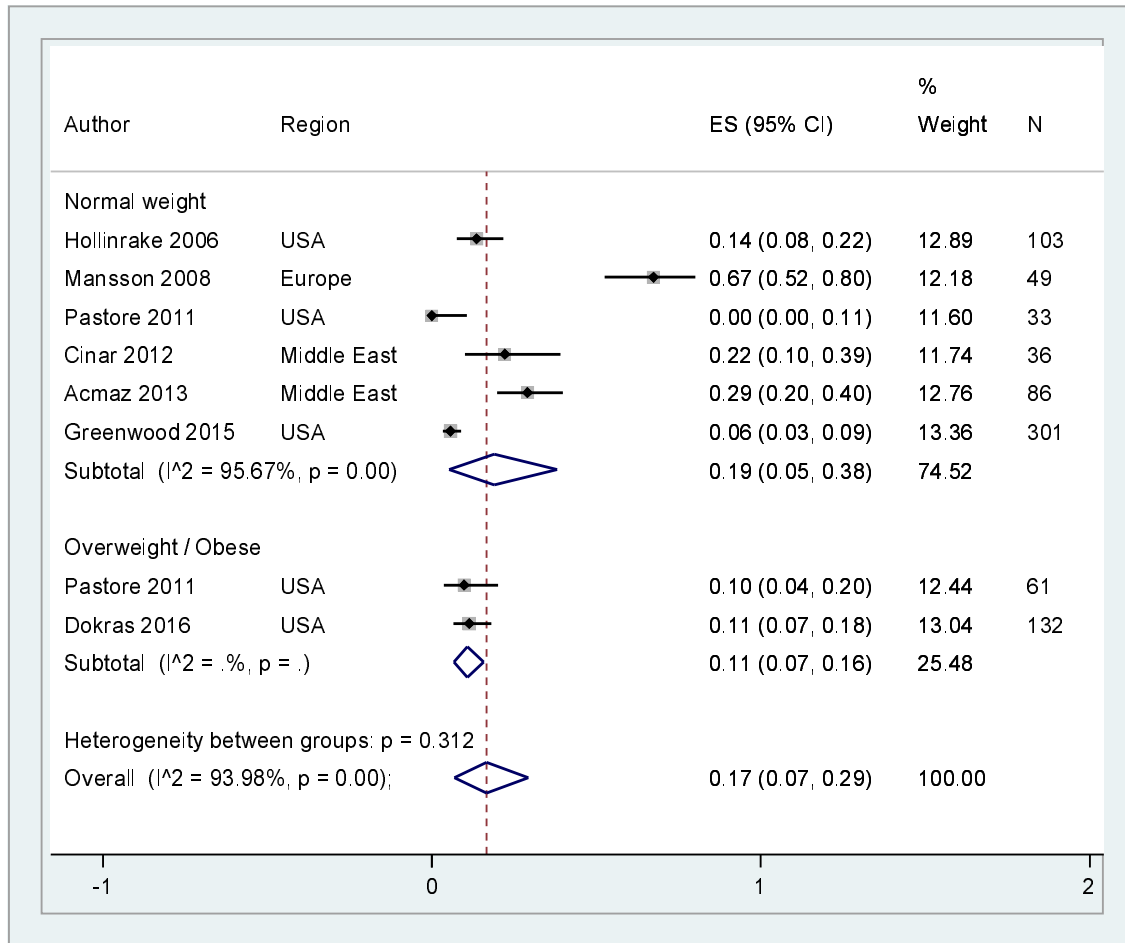


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393 Supplementary a1: Major depression by BMI (sensitivity)

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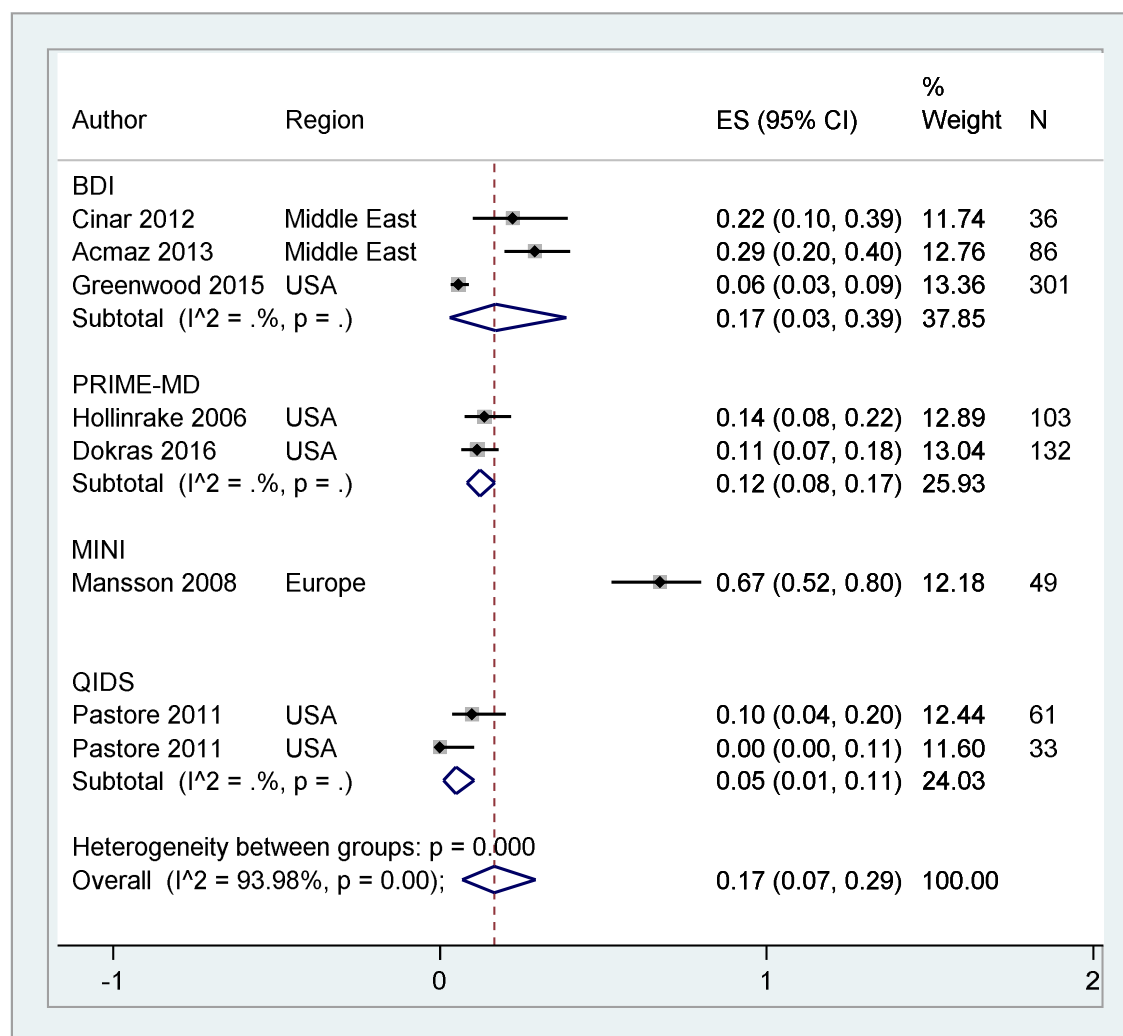
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399 Supplementary a2: Major depression by assessment tool (sensitivity)



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402 QIDS: Quick Inventory of Depressive Symptomatology-Self Report 16,¹

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404 MINI: MINI International Neuropsychiatric Interview,³

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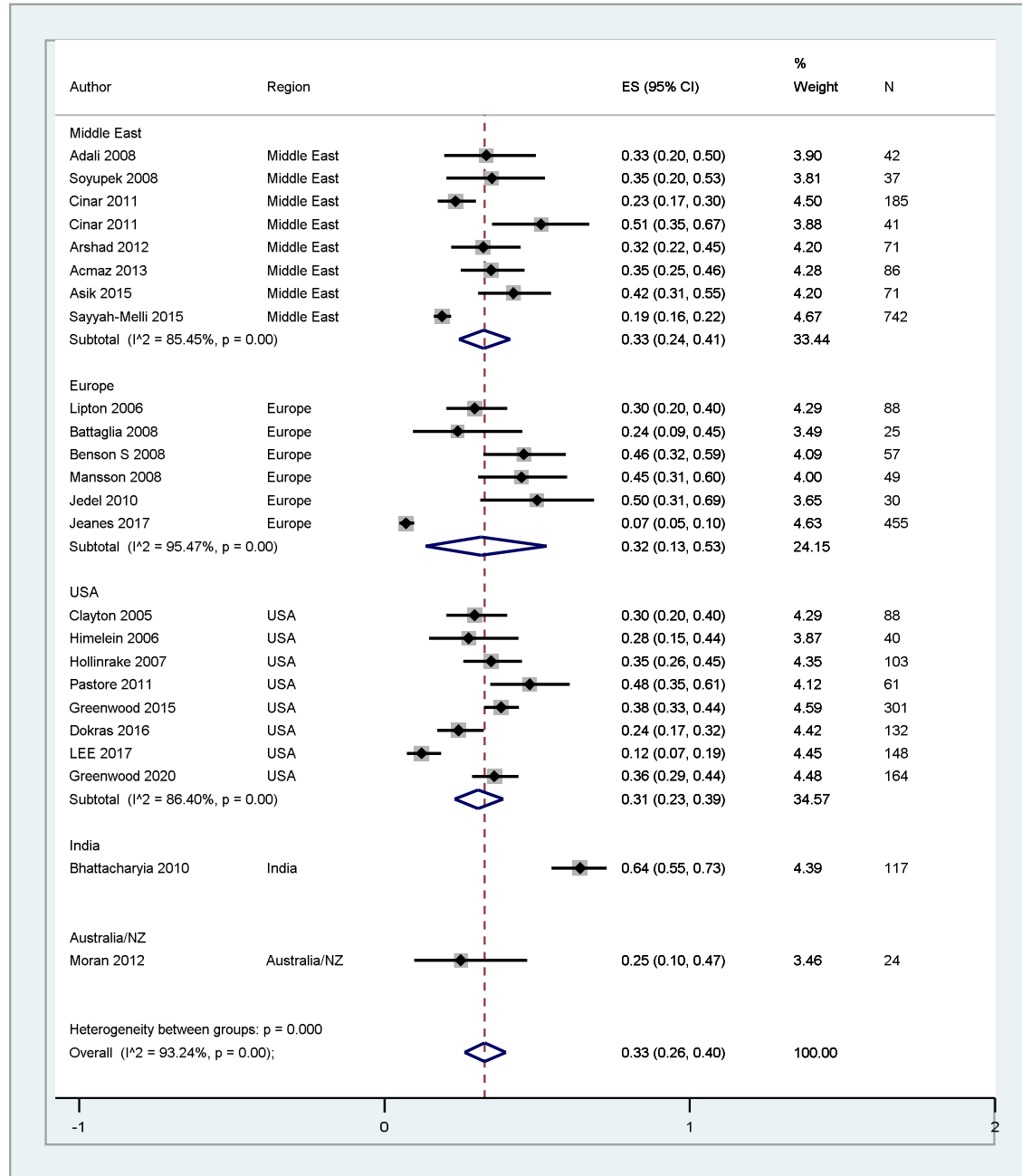
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412 (b) Elevated depression by region

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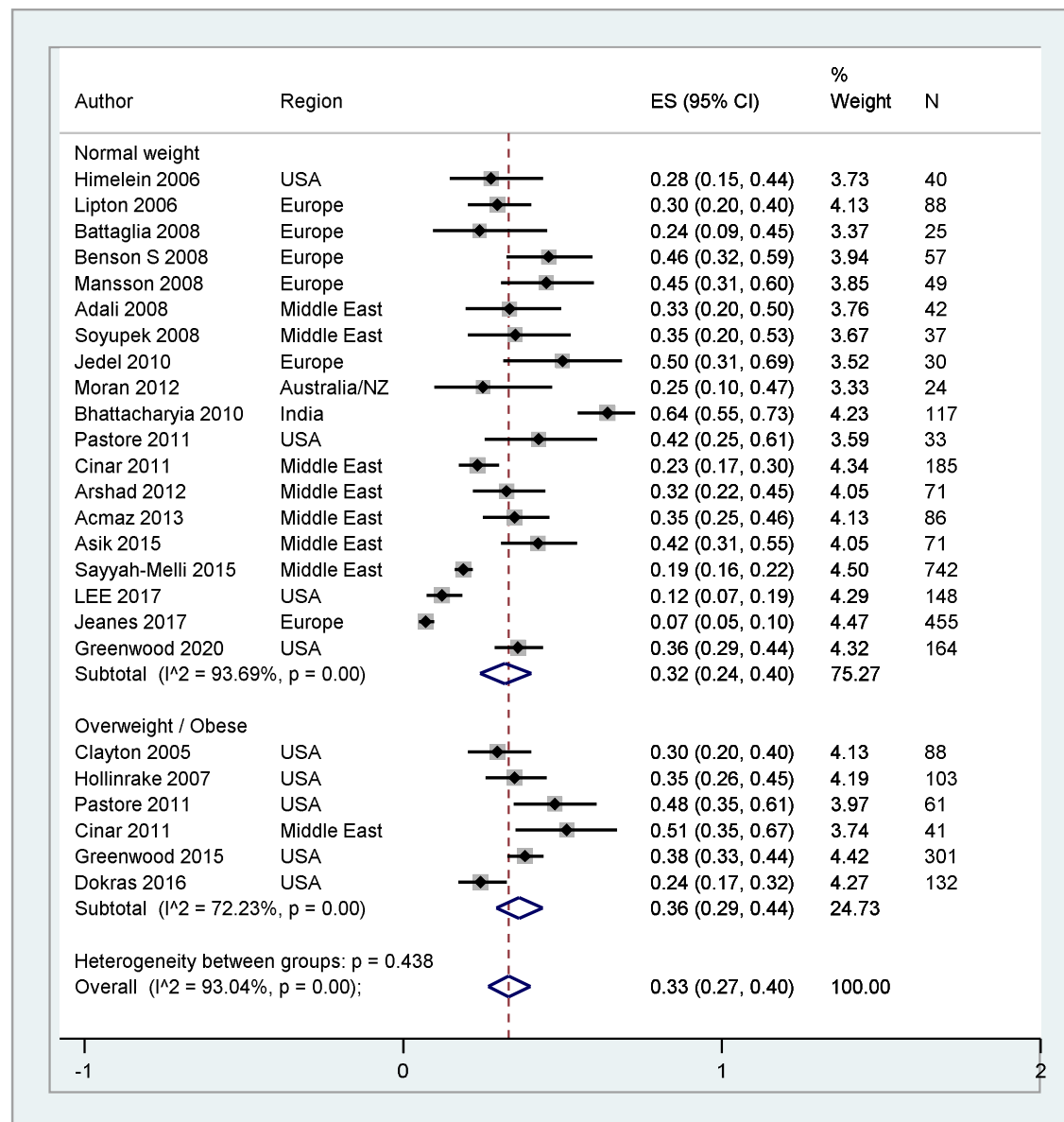
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Supplementary b1: elevated depression by BMI (sensitivity)

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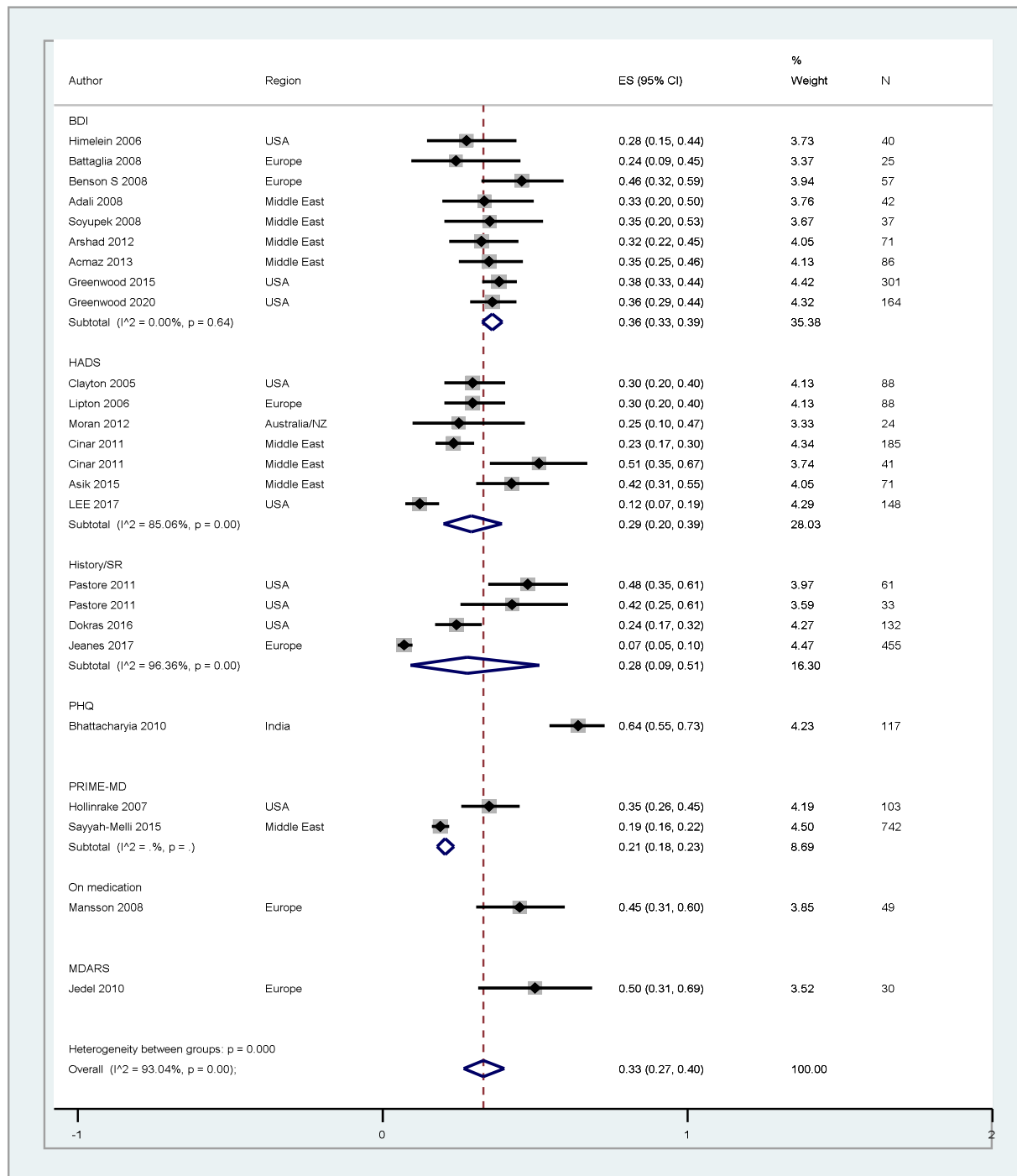
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426 Supplementary b2: elevated depression by assessment tool (sensitivity)

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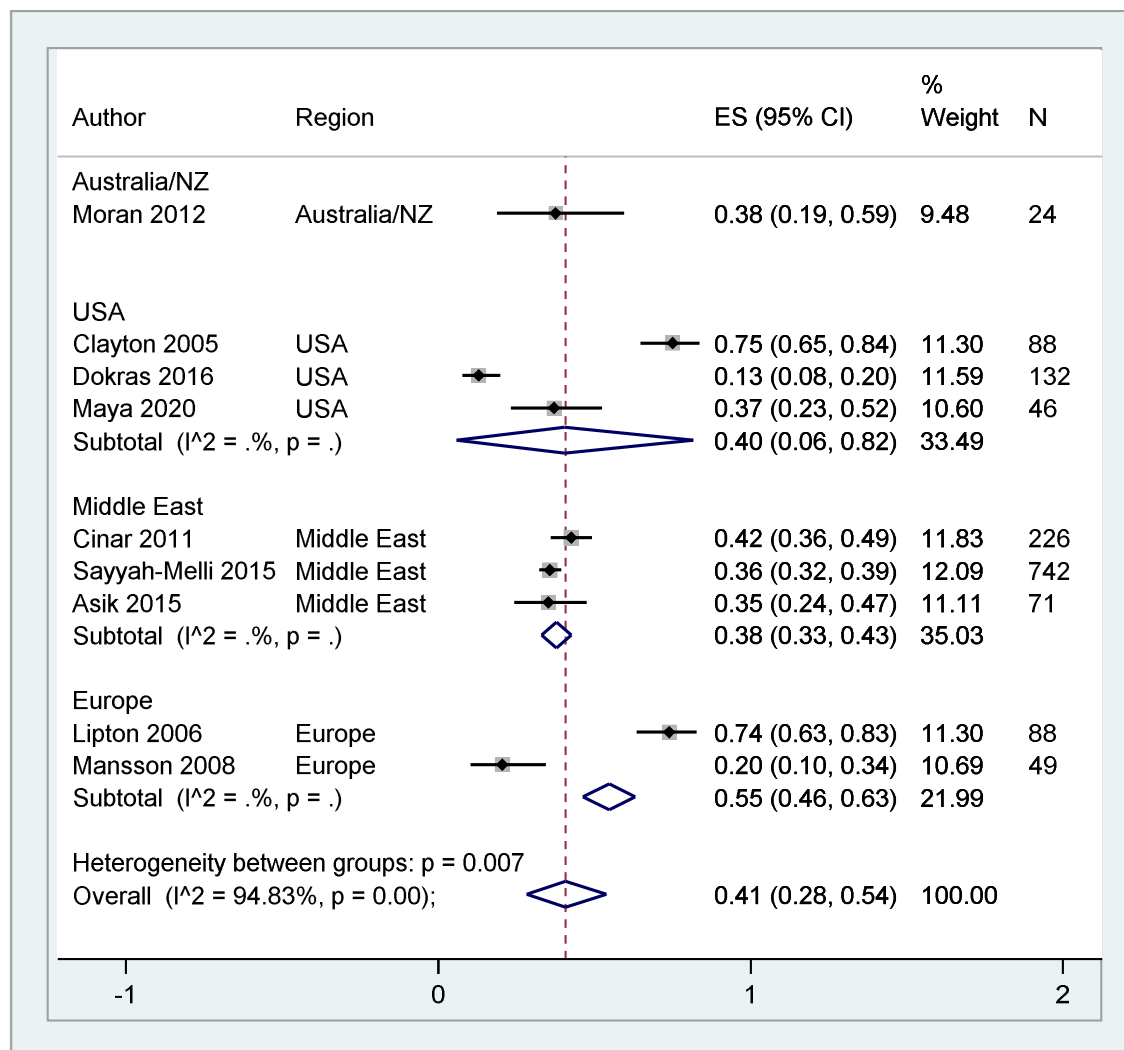
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434 (c) Prevalence of anxiety by region

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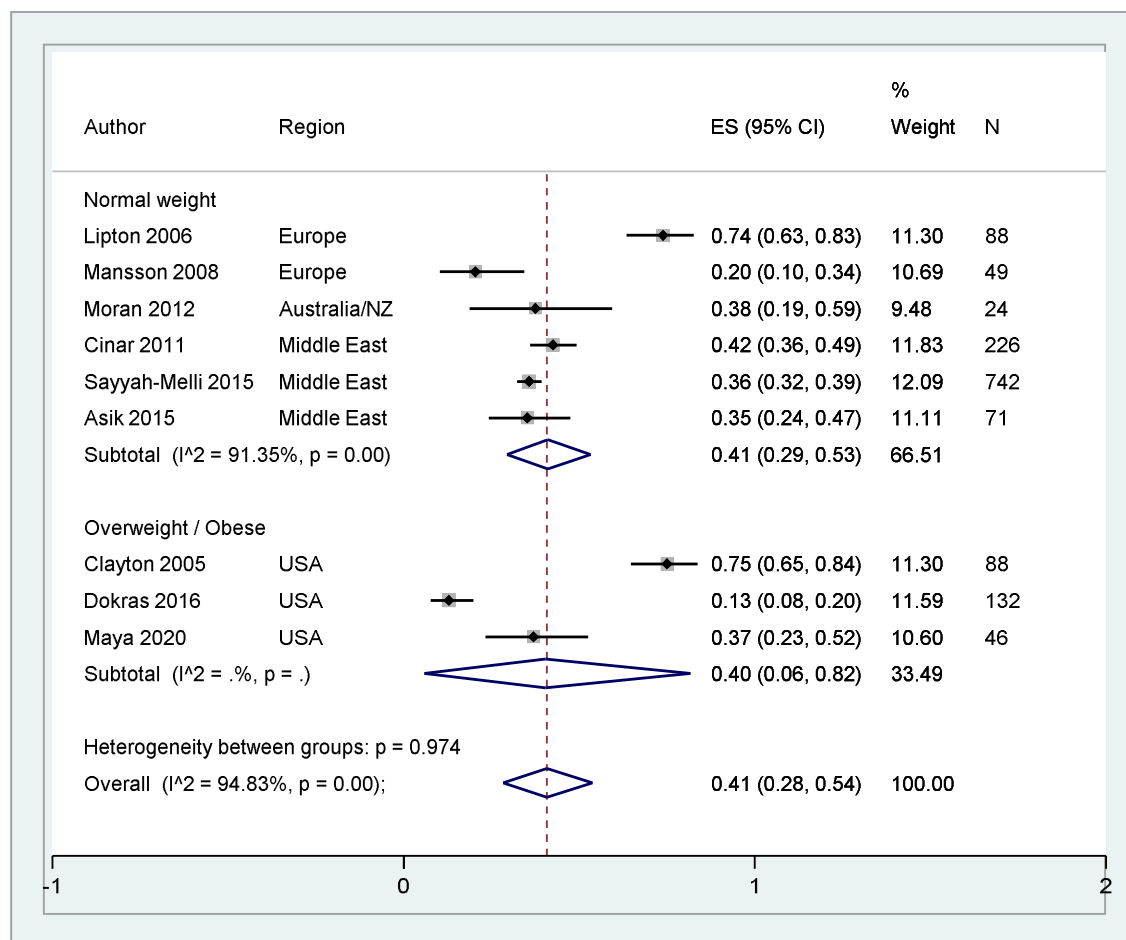
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445 **Supplementary c1: Prevalence of anxiety by BMI (sensitivity)**

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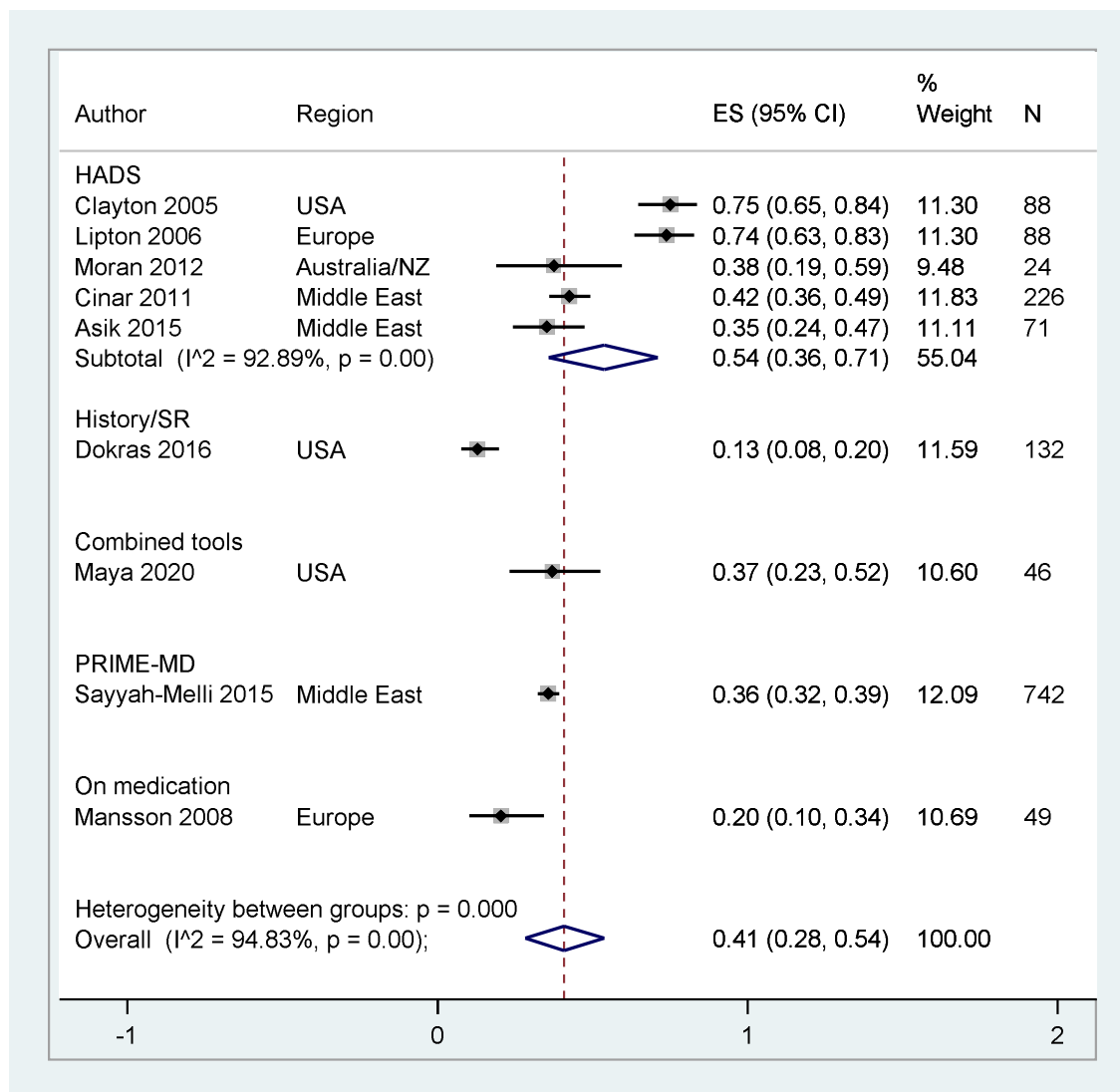


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449 **Supplementary c2: Prevalence of Anxiety by assessment tool (sensitivity)**

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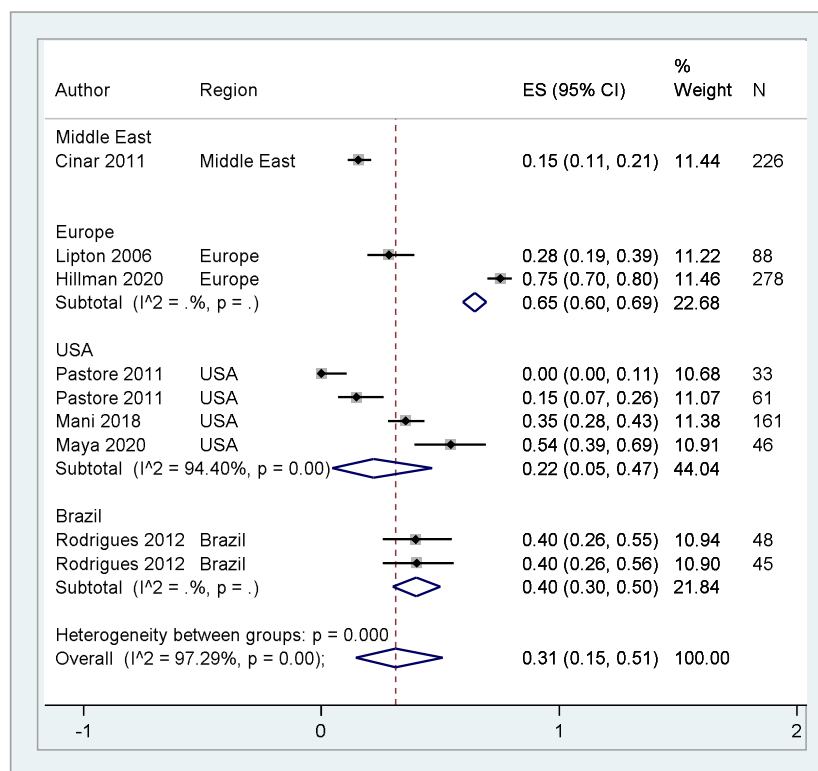
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461 (d) Common mental health by region

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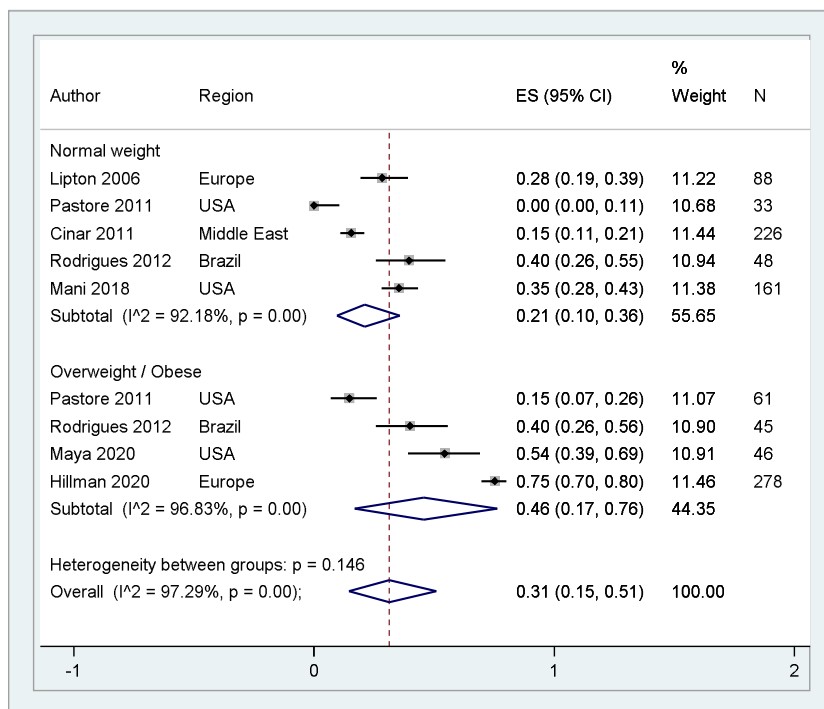


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465 Supplementary d1: Common mental health by BMI (sensitivity)

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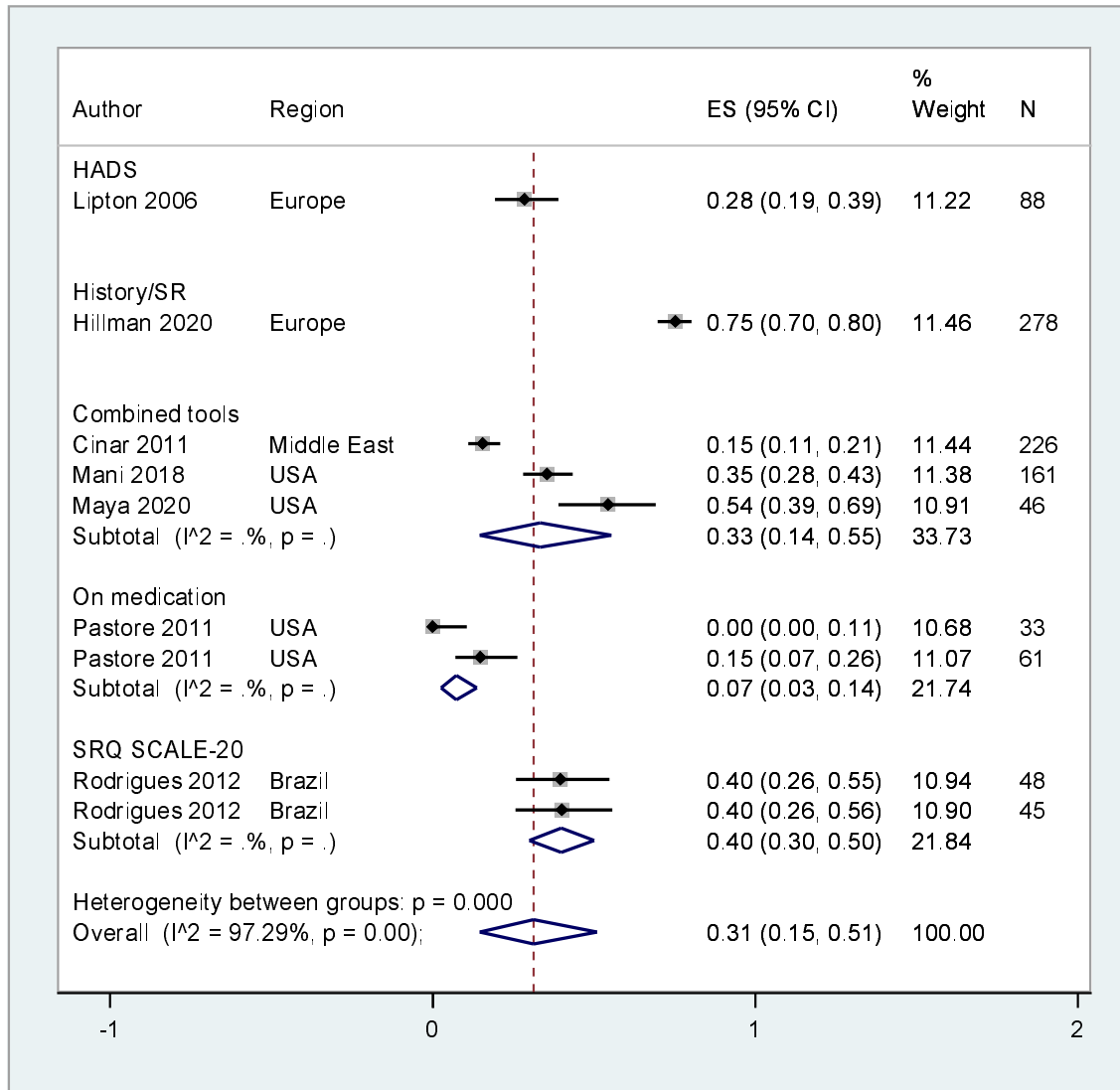


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469 Supplementary d: Common mental health by assessment tool (sensitivity)

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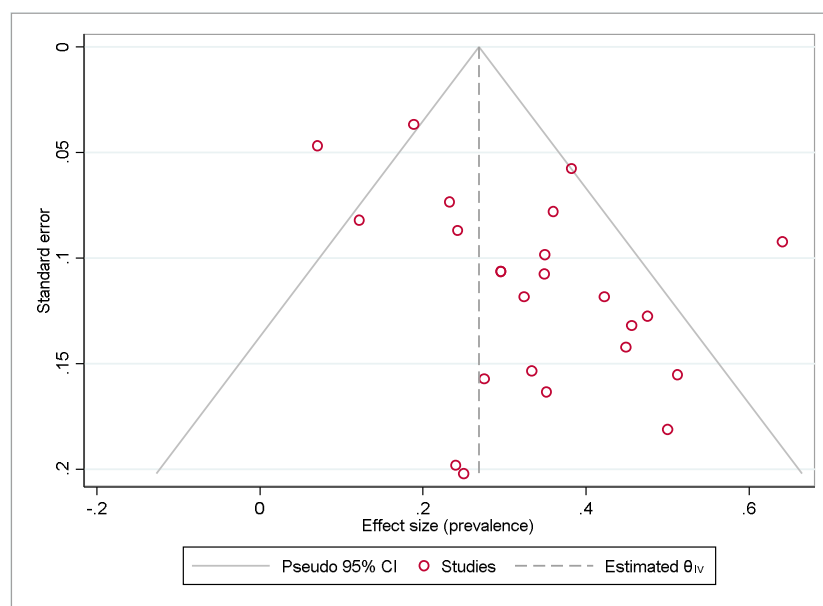


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472 SRQ SCALE-20: Self-reported questionnaire scale-20, 2

473 **FIGURE 2.**

474 (a) Funnel Plot (Prevalence of elevated depression)



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477 Note. The figure visually supports an asymmetrical shaped funnel plot. The no small
478 study effect null hypothesis (H_0 : no small-study effect) may therefore be rejected.

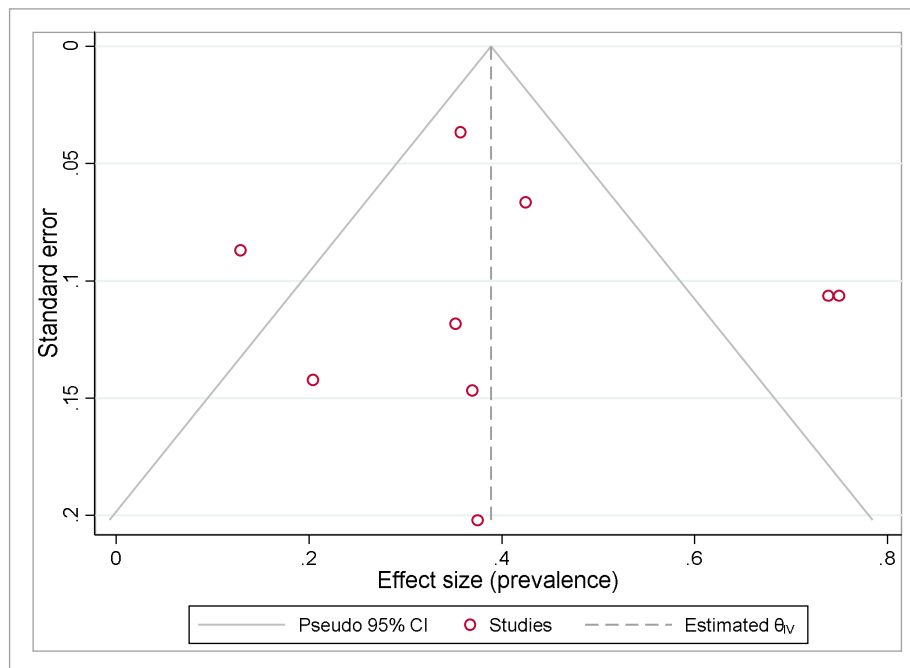
479 Using Egger test of H_0 : no small-study effects, p value = 0.004.

480

481

482 (b) Funnel Plot (Prevalence of anxiety)

483



484

485

486

487 (c)

488 Note. The figure visually supports the symmetry of the effect size distribution. The
489 correlation between effect size and the corresponding standard error was not
490 strong, Egger's test p value = 0.65 supporting no small-study effect (symmetry)
491 hypothesis.

492

493 Discussion

494

495 The meta-analysis identified 11 studies which reported a 20% prevalence of
496 common MH symptoms of anxiety and depression primarily. However, there was
497 substantial heterogeneity across all studies which is reflected by I^2 of 97.29%.
498 Sources contributing to this heterogeneity may include differences in age range,
499 body mass index (BMI), region, study design (including psychiatric assessments

500 used), sample size and methodology, population ethnicity and other characteristics.
501 The narrative synthesis identified several key themes including symptoms of
502 depression and anxiety, as well as diagnosed conditions of eating disorders, bipolar
503 disorder, PTSD and psychotic disorders as demonstrated in **Table x**.

504

505 Despite being the most common reproductive endocrine disorder that spans through
506 the entire reproductive life of a woman, PCOS is often associated with a significant
507 negative patient experience, including delayed diagnosis, that could attribute to
508 exacerbation of symptoms.⁵⁶⁻⁵⁸ Due to the chronic nature of the disease, challenges
509 with persistent treatment adherence is a common issue, but lifestyle changes can be
510 useful to decrease the long-term consequences of the disease. However, once
511 diagnosed, only 19% of women report being aware of all conditions associated with
512 PCOS following discussions with their general practitioner (GP).⁵⁹ The typical
513 symptoms of PCOS such as disturbed menstrual cycles, hirsutism, male-pattern
514 baldness, acne, obesity, and infertility^{12,60-63} could have a further negative impact on
515 patient self-esteem leading to psychological burden. However, narrative accounts
516 point to considerable emphasis on a few symptoms such as infertility by the
517 clinicians, rather than taking a holistic view on all important features of the
518 condition,⁵⁹ such as the long term physical and mental health sequelae, and the
519 effect of diagnosis and treatments on quality of life of the woman. PCOS is also
520 associated with a higher rate of psychiatric disorders.⁵⁶ In keeping with this, our
521 meta-analysis reported 31% prevalence of common MH symptoms and diagnoses in
522 women with PCOS. Similarly, Hillman et al. found an overwhelming 74.9% of women
523 diagnosed with PCOS in their study, to have reported that the condition has
524 adversely affected their MH.⁵⁹ However, despite the considerable negative effect on

525 psychological wellbeing, as many as 34.9% of women in the study chose not to
526 discuss associated MH symptoms with their GP.⁵⁹ An important factor contributing to
527 this may be the ethnic background of the patient. Hillman et al. report that caucasian
528 women in UK diagnosed with PCOS were more likely to discuss their MH with their
529 GP when compared with asian women in UK with PCOS (odds ratio [OR] = 1.66;
530 95% confidence interval [CI] = 1.04 to 2.63).⁵⁹ Clinicians and healthcare providers
531 should better evaluate the needs of BAME communities. This would allow
532 personalised care to be delivered in a culturally sensitive way, which may improve
533 overall patient and clinical outcomes.

534 Around 1 in 6 adults in England meet the diagnostic criteria for common MH
535 disorders.⁶⁴ BAME ethnic groups are more likely to be detained under the Mental
536 Health Act compared to their White British counterparts, and this can be explained
537 by higher rates of mental illness observed and poorer levels of support.⁶⁵ Several
538 studies have explored beliefs about mental illness within BAME communities, and
539 the barriers faced when accessing care. Factors which account for poor presentation
540 rates with mental illness amongst the BAME community include associated stigma,
541 including those with PCOS, poor English language skills, previous negative
542 experiences with healthcare professionals, and being aware of available services.⁶⁶
543 The MH of patients with PCOS may be severely impacted by these issues and
544 reluctance to seek help may further exacerbate pre-existing psychiatric conditions, or
545 limit the recognition of new MH symptoms. Therefore, a thorough assessment of MH
546 illness risk is particularly important in BAME women with PCOS. In agreement, the
547 Royal College of Obstetricians and Gynaecologists (RCOG) recommends the routine
548 screening for psychological symptoms in all women with PCOS.⁶⁷ Yet, our meta-
549 analysis depicts that there is limited number of studies exploring psychological

550 comorbidities in PCOS patients, both in UK and worldwide. Additionally, our meta-
551 analysis was unable to meet most pre-planned outcomes, including the effects of
552 ethnicity and race on MH sequelae in PCOS women, due to the lack of data
553 identified within systematically pooled studies. This was largely due to the non-
554 reporting of patient race and ethnicity in PCOS studies examining patient MH.

555 The two most identified MH conditions amongst PCOS patients in our systematic
556 review were depression and anxiety. Indeed, several existing studies comment on
557 the high risk of both existing and new diagnoses of depression in women with
558 PCOS.^{68,69} Our meta-analysis confirmed these findings, as major depression
559 affected 17% of PCOS patients, and elevated existing depression affected 33%, with
560 a considerably high heterogeneity in both subthemes within and between regions.
561 Existing literature in the field measuring true prevalence of depression in BAME
562 patients with PCOS is contradictory. This meta-analysis could not pool race or
563 ethnicity with the data due to differences in research study designs and clinical
564 practices, as well as the lack of BAME patient representation within studies.
565 Therefore, the true prevalence of elevated existing depression and newly diagnosed
566 major depression amongst BAME PCOS women could not be estimated. However,
567 our narrative synthesis identified certain races/ethnicities to carry higher rates of
568 depression than Caucasian PCOS patients.

569

570 Anxiety disorders include generalised anxiety disorder, panic disorder, separation
571 anxiety disorder, social anxiety disorder and specific phobias⁷⁰ and may impair daily
572 functioning. Women are twice as likely to develop anxiety disorders than men,⁷⁰ with
573 women with PCOS have a significantly higher prevalence of generalised anxiety
574 symptoms than women without PCOS.⁷¹ This study revealed that there are a small

575 number of published studies commenting on anxiety prevalence in women with
576 PCOS. Of these, we identified 41% prevalence of generalised anxiety disorder
577 among women with PCOS, however, heterogeneity was high at 94.83%. This
578 heterogeneity extended to results from different regions suggests lack of difference
579 between geographical locations in prevalence of anxiety amongst women with
580 PCOS. Similar to our depression subtheme analysis, we were unable to establish the
581 prevalence of anxiety amongst different ethnicities and races in women with PCOS
582 due to lack of robust data.

583

584 Eating Disorders (EDs) have high prevalence particularly in young women
585 worldwide.⁷² Body image disturbances and self-esteem are central to the
586 development of common EDs and conditions that may affect body image has been
587 found to contribute to the development of EDs.⁷³ The body dissatisfaction associated
588 with features such as weight gain, hyperandrogenism, hirsutism, acne, and
589 androgenic alopecia often contribute to disturbed body image perception linked with
590 EDs. Existing literature, although informative on the relationship between PCOS and
591 EDs, struggles to report findings on ethnic differences in ED and PCOS. Only two
592 observational studies were identified addressing EDs in BAME PCOS patients,^{74,75}
593 and although they report an increased risk of disordered eating habits in PCOS
594 women compared with the control populations, in agreement, one study found Black,
595 Hispanic and Mixed race women with PCOS to have higher EDE-Q scores (a self-
596 rating questionnaire of the range and severity of ED features) compared with White
597 women⁷⁵ yet the other reported EDs to be similar in the BAME PCOS population.⁷⁵
598 However, discordant methodology and the use of external cohorts from a different
599 country, makes the available evidence to be substandard to draw firm conclusions

600 on the specific prevalence of EDs in BAME women with PCOS, indicating the need
601 for further research to understanding ethnic differences in PCOS and associated
602 EDs.

603

604 Psychotic disorders, such as schizophrenia have been reported to be significantly
605 more prevalent in patients with PCOS in a national study in Sweden.⁷⁶ However,
606 there is a dearth of studies investigating the associations between conditions.
607 Although people from BAME background had been reported to experience a higher
608 prevalence rate of psychosis⁷⁷ compared to the Caucasian population in UK, only
609 one cohort study explored the prevalence of psychotic disorders in Taiwanese PCOS
610 patients,⁷⁸ and reported no statistical difference in prevalence of schizophrenia
611 between the two populations.⁷⁸ This highlights the need for future high quality
612 studies to assess psychotic illnesses relevant to PCOS amongst different ethnicities.

613

614 Bipolar disorder (BD) has been reported to be more prevalent in women with
615 PCOS⁷⁹ compared to the general population and since it carries a 12-fold increased
616 risk of suicide,⁸⁰ it is especially important to examine any causative common
617 mechanistic relations between the two conditions. However, there is a caveat, in that
618 there is a reported high prevalence of menstrual disorders in women receiving
619 treatment for BD.⁸¹ Therefore, accounting for confounding variables such as
620 medication and pre-existing conditions is vital in studies that assess prevalence of
621 MH sequelae, such as BD, in BAME PCOS women. A study from Kashmir Valley⁸²
622 reported BD rates of 2.72% compared to 0.00% in control participants while a
623 nationwide cohort study from Taiwan⁷⁸ found no significant alteration in prevalence
624 of BD in PCOS women compared to a control population. Further research is thus

625 required in order to develop understanding of the prevalence in different populations,
626 as associations cannot be drawn with the limited data available.

627

628 This systematic review had further revealed the heterogeneity of diagnosing
629 psychiatric disorders. Clinicians have adopted 8 different self-report questionnaires
630 in their methodologies highlighted by this review. This limits our study's
631 generalisability as well as any estimate of prevalence of a given psychiatric disorder.

632

633 Overall, although most studies used Rotterdam Criteria in the diagnosis of PCOS,⁵ 4
634 studies adopted the National Institutes of Health Criteria, 1990 to diagnose PCOS,
635 which will bring obvious diagnostic bias.⁸³⁻⁸⁶ Future studies should strive to adapt
636 internationally accepted, standardised criteria for the diagnosis of both PCOS and
637 MH disorder to avoid sampling bias.

638

639 Medical practitioners may not be aware of the impact different cultures may have on
640 patient's reporting of both physical and psychiatric symptoms. In general, members
641 of BAME communities are less likely to engage with MH services or seek help before
642 symptoms severely impact their function.^{87,88} Simkhada et al. found that members of
643 Nepali and Iranian communities report improved MH care and support by healthcare
644 professionals who were trained on different cultural practices.⁸⁹ Furthermore,
645 clinicians in the study stated that a better understanding of different cultures to be
646 beneficial for providing a culturally sensitive service. Therefore, training healthcare
647 professionals in cultural norms within BAME communities is vital to better
648 understand their knowledge and beliefs of MH symptoms, thereby allowing clinicians
649 to provide a holistic, effective treatment approach when working with diverse

650 populations. Cultural competency training may also increase patient trust in clinicians
651 and lead to early discussions of how PCOS may be affecting patients psychosocially
652 within distinct BAME communities. This will therefore aid in early detection and
653 effective and lasting treatment of MH sequelae of PCOS and personalise patient
654 clinical care.

655

656 **Limitations**

657 Several limitations affected our systematic review and meta-analysis findings. Whilst
658 studies listed often identified severe emotional distress amongst PCOS women,
659 some of this was self-reported and therefore subjective, without the presence of a
660 clinical diagnosis. This may mask the true prevalence of common MH conditions,
661 such as depression and anxiety, amongst women with PCOS. A further limitation
662 includes the high heterogeneity observed across all subtheme meta-analyses.
663 Several sources of heterogeneity were behind the variations across estimates.
664 These include the range of scales used, the choice of threshold, although many are
665 equivalent but nonetheless, some disagreement remains and contributes to
666 variations. Other factors such as differences in age range, body mass index (BMI),
667 region, study design, sample size and methodology, population ethnicity and other
668 characteristics, and the established unobserved heterogeneity, all tapping into the
669 differences observed across studies.

670 The cultural differences in reporting MH symptoms across different regions,
671 potentially further impacted by complex interaction with stigma and accepted norms
672 within BAME communities may have specific effects on different populations.
673 Moreover, the difference in reporting MH symptoms and diagnoses is a further
674 limitation of all studies included in this review. Several diverse assessment tools

675 were used in the assessment of risk of key psychiatric disorders, including self-
676 reported symptoms, self-reported use of psychiatric medication and clinical
677 diagnosis, thereby introducing heterogeneity in studies included in this review. This
678 may limit the generalisability of the data obtained, may mask the true prevalence of a
679 given psychiatric disorder due to questionnaires' tendency to screen rather than
680 diagnose.

681

682

683 **Conclusion**

684

685 This study demonstrates the lack of ethnic minority representation in research
686 studies conducted among PCOS patients exploring a possible MH sequelae. To
687 determine if there is a bi-directional relationship between PCOS and MH conditions
688 among ethnic minorities, comprehensive research studies should be designed and
689 conducted as part of a global initiative. A key attribute to the changing needs within
690 women's physical and MH is the migratory patterns that changes the regional and
691 global population. The associated nuances influence both clinical practice and the
692 access to the health-care system, and therefore, regular, and careful scrutiny of
693 contemporary evidence is essential to optimise the overall clinical care offered to
694 these patients. As a result, cultural appropriation based training should be made
695 available and accessible to healthcare professionals within primary and secondary
696 care settings.

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700 **Data sharing statement**

701 All data used within this study has been publicly available. The authors will consider
702 the sharing the dataset gathered upon request.

703

704 **Contributors**

705 GD and DKH developed the systematic review protocol and embedded this within
706 the ELEMI project's evidence synthesis phase. GD, VB, DKH, SR, + SA wrote the
707 first draft of the manuscript. This was furthered by PP, AS, WG, NT, YZ, JS and SR.
708 KM, GD, YZ, VB and JS shared database searches, study selection and extraction
709 for analysis. SA and GD conducted the analysis including the design of the statistical
710 analysis plan. GD, YZ, DKH, PP, KE, WG, NT, YZ, JS, KM, VR, AS, PB, HM, AM,
711 WG and KM critically appraised and finalised the manuscript. All authors approved
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714

715

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727

728 **Supplementary Materials**

729 All supplementary materials associated with this article can be found in the attached

730 Appendices.

731

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