

Endometriosis and physical activity: A narrative review

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1	Endometriosis and Physical Activity: A Narrative Review
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Abstract

Endometriosis is a painful gynecological disorder that affects many women. Constant
treatments and contraception changes affect women looking for solutions to treat and limit the
problems caused by endometriosis. The current narrative review discusses the effects of
physical activity on the management, pain and quality of life in patients with endometriosis.
Findings suggest that body awareness practices such as halta yoga, the Jacobson method, and
progressive muscle relaxation reduce pain and stress and improve the quality of life associated
with endometriosis. It also highlights the current knowledge gap on available evidence and
future research. Moreover, care must be taken when considering patients' physical abilities and
goals and avoiding intense physical activities. The forms of endometriosis and its symptoms
vary from one woman to another, so it is important to carry out studies with various non-
medicinal or surgical techniques. It is important to continue randomized controlled trials to
obtain more data on the benefits of physical activity in women with endometriosis and also to
identify what types of activities could be beneficial to combat pain symptoms and improve the
daily lives of women with endometriosis.

Keywords: Exercise training; pain; quality of life; stress; systematic review

Synopsis

Body awareness practices such as halta yoga, the Jacobson method, and progressive muscle relaxation reduce pain, stress and improve the quality in women with endometriosis.

1. Introduction

- 55 Endometriosis affects 1.5 to 2.5 million or 2- 10% of women of reproductive age [1]. The
- funding allocation for endometriosis is expected to be \$16 million in the NIH budget in 2022.
- 57 The variable and broad symptoms of endometriosis mean that healthcare complicates
- diagnosis, often causing lengthy delays between the onset of symptoms such as pain, abnormal
- 59 menstrual flow, excessive menstrual cramps, and diagnosis [2].
- 60 Endometriosis is a complex gynecological disease with poorly understood mechanisms. The
- main theories utilized to explain the pathogenesis of endometriosis are Sampson's theory, the
- 62 coelomic metaplastic theory, the stem cell theory, the Müllerian remnant theory, and the
- vascular and lymphatic metastasis theory [3].
- 64 Advances in the understanding of endometriosis now include less invasive and
- nonpharmacological treatments [4]. International clinical guidelines suggest focusing on the
- role of physical activity and exercise in managing endometriosis-associated symptoms [5]. The
- 67 inflammation that defines endometriosis sensitizes pelvic organs and leads to chronic pelvic
- pain [6]. This mechanism makes suggests that the anti-inflammatory effects of physical activity
- 69 can likely ameliorate the pain associated with endometriosis [7].
- 70 Some interventions reduce the risk of developing endometriosis [8,9]. However, some authors
- discuss the insufficiency of the existing literature on the effects of physical activity and exercise
- on endometriosis-related symptoms [8].
- 73 Two previous systematic reviews on the effects of physical activity and exercise on
- endometriosis-associated symptoms focused mainly on other complementary and alternative
- 75 treatment options, such as mind–body interventions and acupuncture [9,10]. There is growing
- research interest in endometriosis. We conducted a narrative review to evaluate the benefits of
- physical activity on pain and quality of life in women with endometriosis.
- A narrative review was performed; we presented a non-systematic summation and critical
- analysis of current knowledge on the benefits of physical activity on pain and quality of life in
- 80 women with endometriosis. To refine the scope of our narrative review, we established a set of
- 81 inclusion criteria: articles published up to November 2022 in the English language; original
- 82 scientific papers, clinical trials, meta-analyses, and reviews focusing on the effects of physical
- 83 activity on women with any degree of endometriosis. Physical activity is defined as "any body
- 84 movement produced by skeletal muscles that requires energy expenditure" [11]. Case reports
- or series and letters were excluded. Moreover, articles addressing primary dysmenorrhea were
- 86 excluded as it usually occurs in women without pelvic pathology. PubMed, Scopus, EMBASE,

- Web of Science, ScienceDirect and Cochrane Library were used as electronic databases for
- 88 this research.

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2. The endometriosis

- 90 The diagnosis of endometriosis is complex since the disease takes various forms with different
- 91 symptoms. This often implies a wandering diagnosis, as it takes 5 to 7 years for many women
- 92 to appreciate the medical reasons for the deterioration of their quality-of-life markers.
- 93 The scientific community's attention has recently focused on the genetic mechanisms that may
- 94 be of significance for the increased rates of endometriosis. This hypothesis is supported by
- 95 findings of a 47% contribution from genetic factors to endometriosis, with the remaining 53%
- 96 attributed to unique environmental effects [12]. A recent meta-analysis demonstrated
- 97 significant genetic links between endometriosis and eleven pain conditions, including
- 98 migraine, back pain, multi-localized chronic pain conditions, and inflammatory pathologies
- 99 such as asthma or osteoarthritis [13]. It is likely that osteoarthritis may be associated with
- fluctuations in estrogen and warrants a detailed exploration of genetic susceptibility [13].
- In addition to this role of genetic factors, late pregnancy, early and late menopause, shortened
- menstrual cycles with heavy and prolonged periods [14], malformation of the Müller canal, or
- exposure to diethylstilbestrol in utero [15] appears to be risk factors for endometriosis.
- Some studies suggest that having a lean body mass or low body fat may increase a woman's
- risk for endometriosis [16]. An inverse relationship between endometriosis and body mass
- index (BMI) is commonly acknowledged [17,18], although some studies of limited impact have
- reported otherwise [19]. Reduction in the adipose stem cell population [20], an anorexigenic
- effect through altered hepatic gene expression [21] and promotion of lipid dysfunction and fat
- loss [22] are key mechanisms thought to be involved in the clinically low BMI phenotype
- observed in women with endometriosis. Ferrero et al. [17] [reported that the BMI of 366
- women with endometriosis was significantly lower compared to that of 248 controls and no
- women with endometriosis were obese.

- 114 Endometriosis can be asymptomatic but can be accompanied by disabling pain, for example,
- progressive and chronic pelvic cyclic pain before and during periods (dysmenorrhea) and
- during sex (dyspareunia) [23].
- 117 The most common symptoms of endometriosis are painful sexual intercourse (deep
- dyspareunia), pain before and/or during menstruation (dysmenorrhea), pain when urinating
- (dysuria) and chronic pelvic pain [24]. The symptoms of endometriosis are differentiated by

120 the localization of the lesions. A lesion in the bladder can lead to dysuria resulting in difficulties 121 and pain during urination, urinary infections, and urinary stones [25]. Endometriosis may 122 superficially involve the peritoneum and the abdominal organs or the ovaries [26]. 123

Endometriosis can also cause digestive disorders when it develops in the intestines, colon, and rectum. Digestive endometriosis is a particularly severe form of the disease and accounts for 5 to 10% of endometriosis. In terms of symptoms, there is pain during defecation, alternating diarrhea, constipation, rectorragia, occlusion phenomena with abdominal bloating, gas, and constipation. Irritable bowel syndrome and endometriosis significantly overlap in symptom presentation due to chronic inflammation, thus leading to chronic pelvic pain [27].

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hydrotherapy) [32].

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3. The treatments of endometriosis

The therapeutic management of endometriosis is related to the symptoms and the patient's perspectives. There are cases of superficial endometriosis that do not require special treatment. In a third of cases, superficial lesions will regress after a few months of treatment or without treatment, allowing some women to use complementary medicines to stabilize the symptoms. Treatment is usually offered when symptoms become severe. Hormonal therapy is the first form of treatment that is offered in addition to pain medications [28]. The choice of treatment is influenced by the patient's effectiveness, unfavorable side effects, long-term safety, expenses, and accessibility. Because they interfere with ovulation, most current hormonal treatments are not suitable for endometriosis patients who want to become pregnant [29,30]. Surgery can also be used to remove endometriosis lesions, adhesions, and scar tissue. However, success in reducing pain symptoms and increasing pregnancy rates often depends on the extent of the disease. In addition, lesions may reoccur even after successful eradication, and pelvic floor muscle abnormalities (e.g., tense pelvic muscles, growth on the pelvis) can contribute to chronic pelvic pain [31]. Patients with chronic pelvic pains with endometriosis rarely benefit from hormonal and surgical management. Secondary changes in the pelvis, including the pelvic floor, central nervous system sensitization, and myofascial dysfunction, can continue to generate pain even after treatment [6]. Patients can benefit from physiotherapy and complementary treatments. Physiotherapy for endometriosis should include physical therapy, manual therapy (including visceral therapy), and spa treatments (including balneotherapy and

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Non-medicinal management of endometriosis with acupuncture, osteopathy, and yoga can also improve the quality of life and can be offered in addition to the medical management of endometriosis.

The chronic inflammatory processes in endometriosis can alter the receptivity of the endometrium. Intra-pelvic inflammatory processes affecting the oocyte and embryo can lead to miscarriages [33] and infertility [34].

Women with endometriosis suffer from chronic fatigue that can be explained by severe, regular pain [35]. This fatigue can lead to insomnia, depression, pain, and occupational stress [36]. Pain, physical and mental fatigue, infertility, and the risks associated with endometriosis will affect the social interactions of these women [37]. Women reported that the endometriosis-related experiences that most impacted their lives in a negative way included depression or anxiety (80%), heavy menstrual bleeding (71%), the need to take prescription treatments (57%) or undergo multiple surgeries (49%) or infertility (46%) [38]. These debilitating effects have an immense impact on a patient's quality of life and overall well-being [39], strain social and intimate relationships [40] and lead to losses in productivity, both at home and in the workplace [41].

4. The endometriosis and the benefits of physical activity

Our understanding of the role of physical exercise in endometriosis can benefit from well-controlled studies using validated instruments for evaluation and follow-up in well-defined study groups and well-established exercise treatment protocols. Experimental models of endometriosis permit the characterization of the time course of the disease in order to elucidate whether physical exercise can alter the development of the endometriosis while also making it possible to determine the conditions of physical exercise that can be used in both a preventive and curative manner in endometriosis.

and curative manner in endometriosis.

However, these interventions have been studied mostly in terms of their ability to reduce the risk of developing endometriosis [7,42,43], and so little is known about the effect of PA and exercise on symptom improvement in women with endometriosis [7]. The effect of PA and exercise specifically remained unclear since reviews were limited to a few databases, the search terms were not specified [10], or "PA" and "exercise" were not included as search terms [44]. Studies of interventions involving any type of PA and exercise were eligible for inclusion. PA was defined as "any bodily movement produced by skeletal muscles that requires energy expenditure" [11] and exercise was defined as "PA that is planned, structured, and repetitive for the purpose of conditioning the body" [11], consisting of cardiovascular conditioning, strength and resistance training, and flexibility. Low-intensity exercises are useful treatments for women with endometriosis with or without conventional drug treatments. However, physical activity is differentiated according to the intensity of the effort required. Physical activities that combine breath control, physical postures and meditation (such as Halta Yoga or

Jacobson's progressive relaxation techniques, which consists of voluntarily contracting different muscle groups for periods of 5 to 10 seconds and then releasing the tension) were tested over periods of 6 to 12 weeks and compared with other more traditional physical activities such as muscle strengthening, walking or more intense exercises [44].

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The theory supporting PA and exercise as a beneficial approach involves viewing the skeletal muscles as an endocrine organ, with the contraction of these muscles releasing myokines [45]. These myokines may exert direct effects on the muscle itself or distal organs such as the liver, pancreas, or adipose tissue [45]. Furthermore, exercise increases the production of leucocytes, cortisol, and adrenaline, all of which have potent acute anti-inflammatory effects [46]. Since endometriosis is an estrogen-dependent condition and physical activity may raise levels of sex hormone binding globulin (SHBG), which would lower bioavailable estrogens, it has been hypothesized that physical activity is protective [47]. Additionally, more physical activity lowers insulin resistance and hyperinsulinemia, which have been linked to endometriosis hypotheses [48, 49]. When compared to women who did not regularly exercise, regular exercise was linked to a 40%-80% lower risk of endometriosis [50]. Previous evidence suggests that exercise may be therapeutic in chronic inflammatory processes [15]. Regular physical exercise has shown protective effects in patients with chronic inflammatory disease because it increases systemic levels of anti-inflammatory cytokines [51]. Exercise has previously been described as a possible protective factor for several chronic diseases that involve inflammation, such as inflammatory bowel disease, diabetes mellitus type 2, and breast and colon cancer [52]. Concerning endometriosis, exercise may lead to reduced estrogen levels and frequency of ovulation, increased sex hormone-binding globulin with a subsequent reduction in bio-available estrogen, as well as reduced insulin resistance and hyperinsulinemia. Pain is a debilitating component in the daily lives of women with endometriosis and has been monitored with the EHP-30 questionnaire to show that halta yoga improves the well-being and self-image of women with endometriosis [8]. Likewise, physical activity based on exercises of body consciousness, breathing, stretching and general movements combined with a cognitive approach improved the vitality and physical functioning based on the SF-36 questionnaire [8]. To our knowledge, the study of Ensari et al. [53] was the first to quantify the association

between day-level pain symptoms and exercise in an international sample of women with

endometriosis and to identify habitual weekly exercise frequency as a moderator of this

relationship. They leveraged 90382 days of mHealth self-tracking data from 1009 women with

222 endometriosis to investigate the association between exercise behavior and day-level 223 fluctuations in pain. 224 These participants were likelier to report lower pain scores and smaller increases (or larger 225 decreases) in pain the day after an exercise bout than not having exercised the previous day. In 226 contrast, those who engaged in regular exercise less than twice a week were more likely to 227 experience pain symptoms on days after exercise. According to the physical activity 228 recommendations [54,55], which call for aerobic exercise at least three times per week and 229 muscle-strengthening exercise at least twice per week [56], this is in line with the recommended 230 levels of physical activity. However, the current guidelines make no specific recommendations 231 for endometriosis, and systematic reviews only advise "overall, general exercise" [7,57] 232 because there is insufficient data on the best amount of exercise to relieve endometriosis pain. 233 Only 24.5% (N=202) of the study sample exercised at least three times a week, with the mean 234 weekly frequency of exercise being 1.43/week (SD=1.57, IQR=2.29). This suggests that 235 individuals with endometriosis might be at increased risk of physical inactivity[56], which is a 236 risk factor for various comorbidities [58] and is further linked to exacerbation of chronic pain 237 Ensari et al [53] suggest that their findings of moderate pain in the pelvis as the most frequently 238 reported pain are in line with those from others on endometriosis [61] and various chronic pain 239 conditions[62]. 240 A study by Bergström et al. [63] reported that a 12-month exercise program reduced the 241 intensity of pelvic pain associated with endometriosis, while Carpenter et al. [64] found that 242 physical exercise (6 months) relaxed the muscles of patients with endometriosis to alleviate 243 pain (Table 1). These results are in agreement with the findings of Awad et al. [65] who studied 244 the effects of exercise on pain and posture associated with endometriosis in 20 patients 245 diagnosed with mild or moderate forms of the disease. Their results indicated decreases in the 246 intensity of endometriosis pain after performing exercise and a statistically significant decrease in the thoracic kyphosis angle related to postural kyphosis deformities, having found no 247 correlation between the patient's physical characteristics and intensity of endometriosis pain 248 249 [65]. The results of this study agree with those of a randomized controlled trial indicating that 250 progressive muscular relaxation training was more effective in improving pain, anxiety, 251 depression and fatigue in women with endometriosis regardless of concurrent hormonal 252 therapy [10]. 253 Studies by Carpenter et al. [64] (Table 1) support the beneficial effects of physical exercise in 254 reducing testosterone levels and androgenic symptoms in patients treated with Danazol, a 255 medication used in the treatment of endometrium. Another study reported that physical exercise

mitigated the adverse effects of medications and assisted in the recovery of bone density in women treated with gonadotropin-releasing hormone [63]. Pharmacological therapeutic approaches are available to reduce tissue damage and endometriosis symptoms by suppressing ovulation and suppressing estrogen's stimulative effect on lesions. One of these treatments is with gonadotropin-releasing hormone (GnRH) analog GnRH acts by regulating the anterior pituitary gland and suppressing circulating gonadotropins and sexual hormones [66]. This causes hypoestrogen, resulting in the atrophy of the endometriotic tissue (3). GnRH analogues produce hypoestrogenaemia and increase bone turnover, resulting in substantial bone loss with an imbalance between bone resorption and bone formation [67]. Exercise boosts serotonin levels and increases the firing rate of serotonin neurons in the brain, which stimulates the production of mood-elevating chemicals [9]. Aerobic exercises such as walking increase serotonin levels and muscle tone and improves cardiovascular function [44,63,64]. There is much support for the positive effects of mind-body interventions, such as relaxation and yoga, on overall well-being and quality of life, reducing psychological distress and creating a positive mental state, and decreasing pain intensity [68,69], and pain-associated endometriosis[8,10,44]. Since stress relief should reduce inflammation and pain by activating the HPA axis, it follows that doing so should also reduce painful reactions. Ectopic growth of refluxed endometrium activates a vicious cycle of inflammation characterized by secretion of prostaglandins, growth factors (nerve growth factor-NGF, vascular endothelial growth factor-VEGF), neurotransmitters, and cytokines, into the peritoneal fluid that exacerbates the disease by increasing pain and promoting lesion survival [70]. In a recent prospective study, endometriosis patients who were using positive coping strategies were found to adapt to stress better with less depression and associated pelvic pain [71]. Further, preliminary research found a significant reduction in pain and depression scores when physiotherapy and psychological interventions were carried out in women with endometriosis[72]. Prior research in the field of complementary and integrative medicine has found the beneficial effects of stress-coping strategies, yoga, on chronic pain conditions [10,73], inflammation, immune and brain function, as well as anxiety, depression, and stress levels[74]. Disease processes exacerbated by stress can be interrupted by psychological and stress reduction interventions, including social coping, psychotherapy, exercise, relaxation, meditation, and yoga [73].

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Yoga, classified as a mind-body practice by the WHO [75], covers a myriad of contemplative techniques that use relaxation strategies, including breathing exercises, progressive muscle

relaxation, and relieve stress, helping to integrate the body and mind[76] and alleviates symptoms of asthma and high blood pressure[77,78]. However, research on the various benefits of yoga in the treatment of endometriosis is still scarce, although some studies indicate that 8 to 12 -weeks of yoga reduced pain associated with endometriosis [8,10,44] (**Table 1**).

Women with endometriosis have a high perception of stress, low quality of life, and low levels of cortisol [73,79]. Some studies confirm the relationship between practicing yoga and the reduction of pain and stress associated with improved body awareness and empowerment [80,81]. The practice of yoga involves breathing stabilization, which allows for greater regulation of mental and emotional states, leading to greater insight, better perception of the body, and expanded strategies to reduce pain [82–84], as confirmed in a recent report that practicing relaxation promotes relief from muscle tension and stimulates blood circulation in the muscles and viscera[85].

Yoga reduces autonomic nervous system activity by regulating oxygen intake (i.e., increased respiratory volume) to ease muscle tension and pain. The practice of yoga also regulates psycho–neuro–endocrine structures by balancing the hypothalamic–pituitary–adrenal systems to reorganize the autonomic nervous system [84,86]. As the neuroendocrine systems regulating the stress response are involved in the regulation of mood and emotion[87], mental illnesses such as clinical anxiety and depression are associated with increased expression of stress-induced pro-inflammatory cytokines that stimulate the autonomic nervous system and hypersecretion of corticotropin-releasing hormone (CRH), and increase circulatory cortisol and the production of pro-inflammatory cytokines [88,89].

Studies suggest that pain-associated endometriosis may be related to increased cellular oxidative stress and low cortisol levels, which increase inflammatory responses[90–92].

Physical activity increases levels of sex hormone-binding globulin (SHBG) and reduces the bioavailability of estrogens, which may be protective since endometriosis is an estrogen-dependent disease[93].

5. Conclusions

The main objective of this narrative review was to identify the benefits of physical activity on pain and quality of life in women with endometriosis. An important finding is that body awareness practices such as halta yoga, the Jacobson method, and progressive muscle relaxation reduce pain and stress and improve the quality of life associated with endometriosis. Nevertheless, care must be taken when considering patients' physical abilities and goals and

- 321 avoiding intense physical activities. The forms of endometriosis and its symptoms vary from
- one woman to another, so it is important to carry out studies with various non-medicinal or
- 323 surgical techniques.
- 324 It is important to continue randomized controlled trials to obtain more data on the benefits of
- 325 physical activity in women with endometriosis and also to identify what types of activities
- 326 could be beneficial to combat pain symptoms and improve the daily lives of women with
- endometriosis, as suggested by Mira et al [44].

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- 329 Author contribution: CT, AZ, LF and HZ. participated in the conception and design of the
- study. LF, AZ and SEK were responsible for data collection and statistical analysis. CT, AZ,
- 331 SEK, LF, AS, IL, KW, BK and HZ were responsible for writing and finalization of the
- manuscript. All authors contributed to the manuscript and approved the submitted version. All
- authors have read and agreed to the published version of the manuscript.
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337 6. References

- Zondervan KT, Missmer S, Abrao MS, Einarsson JI, Horne AW, Johnson NP, et al.
- 340 Endometriosis Classification Systems: An International Survey to Map Current Knowledge
- and Uptake. Journal of Minimally Invasive Gynecology 2022;29:716-725.e1.
- 342 https://doi.org/10.1016/j.jmig.2022.01.014.
- 343 [2] Agarwal SK, Chapron C, Giudice LC, Laufer MR, Leyland N, Missmer SA, et al.
- 344 Clinical diagnosis of endometriosis: a call to action. American Journal of Obstetrics &
- 345 Gynecology 2019;220:354.e1-354.e12. https://doi.org/10.1016/j.ajog.2018.12.039.
- 346 [3] Tsamantioti ES, Mahdy H. Endometriosis. StatPearls. Treasure Island, FL; 2022.
- 347 [4] Zondervan KT, Becker CM, Koga K, Missmer SA, Taylor RN, Viganò P.
- Endometriosis. Nature Reviews Disease Primers 2018;4:9. https://doi.org/10.1038/s41572-
- 349 018-0008-5.

- 350 [5] Chapron C, Marcellin L, Borghese B, Santulli P. Rethinking mechanisms, diagnosis
- and management of endometriosis. Nature Reviews Endocrinology 2019;15:666–82.
- 352 [6] Aredo J, Heyrana K, Karp B, Shah J, Stratton P. Relating Chronic Pelvic Pain and
- 353 Endometriosis to Signs of Sensitization and Myofascial Pain and Dysfunction. Seminars in
- 354 Reproductive Medicine 2017;35. https://doi.org/10.1055/s-0036-1597123.
- 355 [7] Bonocher CM, Montenegro ML, Rosa e Silva JC, Ferriani RA, Meola J. Endometriosis
- and physical exercises: a systematic review. Reproductive Biology and Endocrinology
- 357 2014;12:4. https://doi.org/10.1186/1477-7827-12-4.
- 358 [8] Tennfjord MK, Gabrielsen R, Tellum T. Effect of physical activity and exercise on
- 359 endometriosis-associated symptoms: a systematic review. BMC Women's Health
- 360 2021;21:355. https://doi.org/10.1186/s12905-021-01500-4.
- 361 [9] Zhao L, Wu H, Zhou X, Wang Q, Zhu W, Chen J. Effects of progressive muscular
- relaxation training on anxiety, depression and quality of life of endometriosis patients under
- 363 gonadotrophin-releasing hormone agonist therapy. European Journal of Obstetrics &
- 364 Gynecology and Reproductive Biology 2012;162:211–5.
- 365 https://doi.org/10.1016/j.ejogrb.2012.02.029.
- 366 [10] Evans S, Fernandez S, Olive L, Payne LA, Mikocka-Walus A. Psychological and mind-
- 367 body interventions for endometriosis: A systematic review. Journal of Psychosomatic Research
- 368 2019;124:109756. https://doi.org/10.1016/j.jpsychores.2019.109756.
- 369 [11] Caspersen CJ, Powell KE, Christenson GM. Physical activity, exercise, and physical
- fitness: definitions and distinctions for health-related research. Public Health Rep 100(2):126-
- 371 131 1985;100.
- 372 [12] Saha R, Pettersson HJ, Svedberg P, Olovsson M, Bergqvist A, Marions L, et al.
- 373 Heritability of endometriosis. Fertility and Sterility 2015;104:947–52.
- 374 https://doi.org/10.1016/j.fertnstert.2015.06.035.
- Rahmioglu N, Mortlock S, Ghiasi M, Møller PL, Stefansdottir L, Galarneau G, et al.
- 376 The genetic basis of endometriosis and comorbidity with other pain and inflammatory
- 377 conditions. Nature Genetics 2023;55:423–36. https://doi.org/10.1038/s41588-023-01323-z.
- 378 [14] Medical management of endometriosis. International Journal of Gynecology &
- 379 Obstetrics 2000;71:183–96. https://doi.org/10.1016/S0020-7292(00)80034-X.
- 380 [15] Petersen AMW, Pedersen BK. The anti-inflammatory effect of exercise. Journal of
- 381 Applied Physiology 2005;98:1154–62. https://doi.org/10.1152/japplphysiol.00164.2004.

- 382 [16] Farland L, Missmer S, Bijon A, Gusto G, Gelot A, Clavel-Chapelon F, et al.
- Associations among body size across the life course, adult height and endometriosis. Human
- Reproduction (Oxford, England) 2017;32:1–11. https://doi.org/10.1093/humrep/dex207.
- Ferrero S, Anserini P, Remorgida V, Ragni N. Body mass index in endometriosis.
- European Journal of Obstetrics & Gynecology and Reproductive Biology 2005;121:94–8.
- 387 https://doi.org/10.1016/j.ejogrb.2004.11.019.
- 388 [18] Liu Y, Zhang W. Association between body mass index and endometriosis risk: a meta-
- analysis. Oncotarget; Vol 8, No 29 2017.
- Tang Y, Zhao M, Lin L, Gao Y, Chen GQ, Chen S, et al. Is body mass index associated
- with the incidence of endometriosis and the severity of dysmenorrhoea: a case–control study
- in China? BMJ Open 2020;10. https://doi.org/10.1136/bmjopen-2020-037095.
- 393 [20] Zolbin MM, Mamillapalli R, Nematian SE, Goetz TG, Taylor HS. Adipocyte alterations
- 394 in endometriosis: reduced numbers of stem cells and microRNA induced alterations in
- adipocyte metabolic gene expression. Reproductive Biology and Endocrinology 2019;17:36.
- 396 https://doi.org/10.1186/s12958-019-0480-0.
- 397 [21] Goetz TG, Mamillapalli R, Taylor HS. Low Body Mass Index in Endometriosis Is
- 398 Promoted by Hepatic Metabolic Gene Dysregulation in Micel. Biology of Reproduction
- 399 2016;95. https://doi.org/10.1095/biolreprod.116.142877.
- 400 [22] Dutta M, Anitha M, Smith PB, Chiaro CR, Maan M, Chaudhury K, et al. Metabolomics
- 401 Reveals Altered Lipid Metabolism in a Mouse Model of Endometriosis. J Proteome Res
- 402 2016;15:2626–33. https://doi.org/10.1021/acs.jproteome.6b00197.
- 403 [23] Pugsley Z, Ballard K. Management of endometriosis in general practice: The pathway
- 404 to diagnosis. The British Journal of General Practice: The Journal of the Royal College of
- 405 General Practitioners 2007;57:470–6.
- 406 [24] Falcone T, Lebovic DI. Clinical Management of Endometriosis. Obstetrics &
- 407 Gynecology 2011;118.
- 408 [25] Maccagnano C, Pellucchi F, Rocchini L, Ghezzi M, Scattoni V, Montorsi F, et al.
- 409 Ureteral Endometriosis: Proposal for a Diagnostic and Therapeutic Algorithm with a Review
- of the Literature. Urologia Internationalis 2013;91:1–9. https://doi.org/10.1159/000345140.
- 411 [26] Ferrero S, Vellone VG, Barra F. Pathophysiology of pain in patients with peritoneal
- 412 endometriosis. Annals of Translational Medicine; Vol 7, Supplement 1 (March 27, 2019):
- 413 Annals of Translational Medicine 2019.
- 414 [27] DiVasta AD, Zimmerman LA, Vitonis AF, Fadayomi AB, Missmer SA. Overlap
- 415 Between Irritable Bowel Syndrome Diagnosis and Endometriosis in Adolescents. Clinical

- 416 Gastroenterology and Hepatology 2021;19:528-537.e1.
- 417 https://doi.org/10.1016/j.cgh.2020.03.014.
- 418 [28] Vannuccini S, Clemenza S, Rossi M, Petraglia F. Hormonal treatments for
- 419 endometriosis: The endocrine background. Reviews in Endocrine and Metabolic Disorders
- 420 2022;23:333–55. https://doi.org/10.1007/s11154-021-09666-w.
- 421 [29] Carey ET, Till SR, As-Sanie S. Pharmacological Management of Chronic Pelvic Pain
- 422 in Women. Drugs 2017;77:285–301. https://doi.org/10.1007/s40265-016-0687-8.
- 423 [30] Harrison JE, Weber S, Jakob R, Chute CG. ICD-11: an international classification of
- 424 diseases for the twenty-first century. BMC Medical Informatics and Decision Making
- 425 2021;21:206. https://doi.org/10.1186/s12911-021-01534-6.
- 426 [31] Zondervan KT, Becker CM, Missmer SA. Endometriosis. New England Journal of
- 427 Medicine 2020;382:1244–56. https://doi.org/10.1056/NEJMra1810764.
- 428 [32] Wójcik M, Szczepaniak R, Placek K. Physiotherapy Management in Endometriosis.
- 429 International Journal of Environmental Research and Public Health 2022;19.
- 430 https://doi.org/10.3390/ijerph192316148.
- 431 [33] Pirtea P, Cicinelli E, Nola RD, Ziegler D de, Ayoubi JM. Endometrial causes of
- 432 recurrent pregnancy losses: endometriosis, adenomyosis, and chronic endometritis. Fertility
- 433 and Sterility 2021;115:546–60. https://doi.org/10.1016/j.fertnstert.2020.12.010.
- 434 [34] Missmer S, Hankinson S, Spiegelman D, Barbieri R, Marshall L, Hunter D. Incidence
- 435 of Laparoscopically Confirmed Endometriosis by Demographic, Anthropometric, and
- 436 Lifestyle Factors. American Journal of Epidemiology 2004;160:784–96.
- 437 https://doi.org/10.1093/aje/kwh275.
- 438 [35] Soliman A, Rahal Y, Robert C, Defoy I, Nisbet P, Leyland N, et al. Impact of
- 439 Endometriosis on Fatigue and Productivity Impairment in a Cross-Sectional Survey of
- 440 Canadian Women. Journal of Obstetrics and Gynaecology Canada 2020;43.
- 441 https://doi.org/10.1016/j.jogc.2020.06.022.
- Ramin-Wright A, Kohl Schwartz A, Geraedts K, Rauchfuss M, Wölfler M, Haeberlin
- 443 F, et al. Fatigue A symptom in endometriosis. Human Reproduction (Oxford, England)
- 444 2018;33. https://doi.org/10.1093/humrep/dey115.
- 445 [37] Bourdel N, Comptour A, Chauvet P, Canis M. Douleurs et endométriose. Douleurs :
- 446 Évaluation Diagnostic Traitement 2020;21:3–10.
- 447 https://doi.org/10.1016/j.douler.2019.11.001.
- 448 [38] Missmer SA, Tu F, Soliman AM, Chiuve S, Cross S, Eichner S, et al. Impact of
- endometriosis on women's life decisions and goal attainment: a cross-sectional survey of

- 450 members of an online patient community. BMJ Open 2022;12.
- 451 https://doi.org/10.1136/bmjopen-2021-052765.
- 452 [39] Gallagher JS, DiVasta AD, Vitonis AF, Sarda V, Laufer MR, Missmer SA. The Impact
- 453 of Endometriosis on Quality of Life in Adolescents. Journal of Adolescent Health
- 454 2018;63:766–72. https://doi.org/10.1016/j.jadohealth.2018.06.027.
- 455 [40] Culley L, Law C, Hudson N, Mitchell H, Denny E, Raine-Fenning N. A qualitative
- 456 study of the impact of endometriosis on male partners. Human Reproduction (Oxford, England)
- 457 2017;32:1–7. https://doi.org/10.1093/humrep/dex221.
- 458 [41] Soliman A, Coyne K, Gries K, Castelli-Haley J, Snabes M, Surrey E. The Effect of
- Endometriosis Symptoms on Absenteeism and Presenteeism in the Workplace and at Home.
- 460 Journal of Managed Care & Specialty Pharmacy 2017;23:745-54.
- 461 https://doi.org/10.18553/jmcp.2017.23.7.745.
- 462 [42] Shafrir AL, Farland LV, Shah DK, Harris HR, Kvaskoff M, Zondervan K, et al. Risk
- 463 for and consequences of endometriosis: A critical epidemiologic review. Best Practice &
- 464 Research Clinical Obstetrics & Gynaecology 2018;51:1–15.
- 465 https://doi.org/10.1016/j.bpobgyn.2018.06.001.
- 466 [43] Ricci E, Viganò P, Cipriani S, Chiaffarino F, Bianchi S, Rebonato G, et al. Physical
- activity and endometriosis risk in women with infertility or pain: Systematic review and meta-
- analysis. Medicine 2016;95.
- 469 [44] Mira TAA, Buen MM, Borges MG, Yela DA, Benetti-Pinto CL. Systematic review and
- 470 meta-analysis of complementary treatments for women with symptomatic endometriosis.
- 471 International Journal of Gynecology & Obstetrics 2018;143:2–9.
- 472 https://doi.org/10.1002/ijgo.12576.
- 473 [45] Brandt C, Pedersen BK. The Role of Exercise-Induced Myokines in Muscle
- 474 Homeostasis and the Defense against Chronic Diseases. Journal of Biomedicine and
- 475 Biotechnology 2010;2010:520258. https://doi.org/10.1155/2010/520258.
- 476 [46] Nimmo MA, Leggate M, Viana JL, King JA. The effect of physical activity on
- 477 mediators of inflammation. Diabetes, Obesity and Metabolism 2013;15:51-60.
- 478 https://doi.org/10.1111/dom.12156.
- 479 [47] Verkasalo PK, Thomas HV, Appleby PN, Davey GK, Key TJ. Circulating levels of sex
- 480 hormones and their relation to risk factors for breast cancer: a cross-sectional study in 1092
- pre- and postmenopausal women (United Kingdom). Cancer Causes & Control 2001;12:47–
- 482 59. https://doi.org/10.1023/A:1008929714862.

- 483 [48] Stoll B. Adiposity as a risk determinant for postmenopausal breast cancer. International
- 484 Journal of Obesity 2000;24:527–33. https://doi.org/10.1038/sj.ijo.0801247.
- 485 [49] Kaaks R. Nutrition, hormones, and breast cancer: Is insulin the missing link? Cancer
- 486 Causes & Control 1996;7:605–25. https://doi.org/10.1007/BF00051703.
- 487 [50] Heilier J-F, Donnez J, Nackers F, Rousseau R, Verougstraete V, Rosenkranz K, et al.
- 488 Environmental and host-associated risk factors in endometriosis and deep endometriotic
- 489 nodules: A matched case-control study. Environmental Research 2007;103:121-9.
- 490 https://doi.org/10.1016/j.envres.2006.04.004.
- 491 [51] Wu M-H, Shoji Y, Chuang P-C, Tsai S-J. Endometriosis: disease pathophysiology and
- 492 the role of prostaglandins. Expert Reviews in Molecular Medicine 2007;9:1-20.
- 493 https://doi.org/10.1017/S146239940700021X.
- 494 [52] Friedenreich CM, Orenstein MR. Physical Activity and Cancer Prevention: Etiologic
- 495 Evidence and Biological Mechanisms. The Journal of Nutrition 2002;132:3456S-3464S.
- 496 https://doi.org/10.1093/jn/132.11.3456S.
- 497 [53] Ipek Ensari, Sharon Lipsky-Gorman, Emma N Horan, Suzanne Bakken, Noémie
- 498 Elhadad. Associations between physical exercise patterns and pain symptoms in individuals
- 499 with endometriosis: a cross-sectional mHealth-based investigation. BMJ Open
- 500 2022;12:e059280. https://doi.org/10.1136/bmjopen-2021-059280.
- 501 [54] DiPietro L, Buchner DM, Marquez DX, Pate RR, Pescatello LS, Whitt-Glover MC.
- New scientific basis for the 2018 U.S. Physical Activity Guidelines. Journal of Sport and
- 503 Health Science 2019;8:197–200. https://doi.org/10.1016/j.jshs.2019.03.007.
- 504 [55] Bull FC, Al-Ansari SS, Biddle S, Borodulin K, Buman MP, Cardon G, et al. World
- Health Organization 2020 guidelines on physical activity and sedentary behaviour. British
- Journal of Sports Medicine 2020;54:1451–62. https://doi.org/10.1136/bjsports-2020-102955.
- 507 [56] Piercy K, Troiano R, Ballard R, Carlson S, Fulton J, Galuska D, et al. The Physical
- 508 Activity Guidelines for Americans. JAMA 2018;320.
- 509 https://doi.org/10.1001/jama.2018.14854.
- 510 [57] Armour M, Ee, CC, Naidoo, D, Ayati, Z, Chalmers, KJ, Steel, KA, de Manincor, MJ,
- Delshad E. Exercise for dysmenorrhoea. Cochrane Database of Systematic Reviews 2019.
- 512 https://doi.org/10.1002/14651858.CD004142.pub4.
- 513 [58] KATZMARZYK PT, POWELL KE, JAKICIC JM, TROIANO RP, PIERCY K,
- TENNANT B, et al. Sedentary Behavior and Health: Update from the 2018 Physical Activity
- 515 Guidelines Advisory Committee. Medicine & Science in Sports & Exercise 2019;51.

- 516 [59] Basen-Engquist K, Scruggs S, Jhingran A, Bodurka D, Lu K, Ramondetta L, et al.
- 517 Physical Activity and Obesity in Endometrial Cancer Survivors: Associations with Pain,
- 518 Fatigue, and Physical Functioning. American Journal of Obstetrics and Gynecology
- 519 2009;200:288.e1-8. https://doi.org/10.1016/j.ajog.2008.10.010.
- 520 [60] Dansie EJ, Turk DC, Martin KR, Van Domelen DR, Patel KV. Association of Chronic
- Widespread Pain With Objectively Measured Physical Activity in Adults: Findings From the
- National Health and Nutrition Examination Survey. The Journal of Pain 2014;15:507–15.
- 523 https://doi.org/10.1016/j.jpain.2014.01.489.
- 524 [61] Warzecha D, Szymusik I, Wielgos M, Pietrzak B. The Impact of Endometriosis on the
- Quality of Life and the Incidence of Depression—A Cohort Study. International Journal of
- Environmental Research and Public Health 2020;17. https://doi.org/10.3390/ijerph17103641.
- 527 [62] Bouhassira D, Lantéri-Minet M, Attal N, Laurent B, Touboul C. Prevalence of chronic
- pain with neuropathic characteristics in the general population. PAIN 2008;136:380-7.
- 529 https://doi.org/10.1016/j.pain.2007.08.013.
- 530 [63] Bergström I, Freyschuss B, Jacobsson H, Landgren B-M. The effect of physical training
- on bone mineral density in women with endometriosis treated with GnRH analogs: a pilot
- 532 study. Acta Obstetricia et Gynecologica Scandinavica 2005;84:380–3.
- 533 https://doi.org/10.1111/j.0001-6349.2005.00558.x.
- 534 [64] Carpenter SE, Tjaden B, Rock JA, Kimball A. The effect of regular exercise on women
- receiving danazol for treatment of endometriosis. International Journal of Gynecology &
- 536 Obstetrics 1995;49:299–304. https://doi.org/10.1016/0020-7292(95)02359-K.
- 537 [65] Awad E, Hamada H, Yousef A, Abbas R. Efficacy of exercise on pelvic pain and
- 538 posture associated with endometriosis: Within subject design. Journal of Physical Therapy
- 539 Science 2017;29:2112–5. https://doi.org/10.1589/jpts.29.2112.
- 540 [66] Pierce SJ, Gazvani MR, Farquharson RG. Long-term use of gonadotropin-releasing
- hormone analogs and hormone replacement therapy in the management of endometriosis: a
- 542 randomized trial with a 6-year follow-up. Fertility and Sterility 2000;74:964-8.
- 543 https://doi.org/10.1016/S0015-0282(00)01537-5.
- 544 [67] Cheng C-H, Chen L-R, Chen K-H. Osteoporosis Due to Hormone Imbalance: An
- 545 Overview of the Effects of Estrogen Deficiency and Glucocorticoid Overuse on Bone
- 546 Turnover. International Journal of Molecular Sciences 2022;23.
- 547 https://doi.org/10.3390/ijms23031376.
- 548 [68] Gautam S, Tolahunase M, Kumar U, Dada R. Impact of yoga based mind-body
- 549 intervention on systemic inflammatory markers and co-morbid depression in active

- 550 Rheumatoid arthritis patients: A randomized controlled trial. Restorative Neurology and
- 551 Neuroscience 2019;37:41–59. https://doi.org/10.3233/RNN-180875.
- 552 [69] Zautra A, Davis M, Reich J, Nicassario P, Tennen H, Finan P, et al. Comparison of
- 553 Cognitive Behavioral and Mindfulness Meditation Interventions on Adaptation to Rheumatoid
- Arthritis for Patients With and Without History of Recurrent Depression. Journal of Consulting
- and Clinical Psychology 2008;76:408–21. https://doi.org/10.1037/0022-006X.76.3.408.
- 556 [70] Coxon L, Horne AW, Vincent K. Pathophysiology of endometriosis-associated pain: A
- review of pelvic and central nervous system mechanisms. Best Practice & Research Clinical
- 558 Obstetrics & Gynaecology 2018;51:53–67. https://doi.org/10.1016/j.bpobgyn.2018.01.014.
- 559 [71] Donatti L, Ramos DG, Andres M de P, Passman LJ, Podgaec S. Patients with
- endometriosis using positive coping strategies have less depression, stress and pelvic pain.
- 561 Einstein (São Paulo) 2017;15.
- 562 [72] Friggi Sebe Petrelluzzi K, Garcia MC, Petta CA, Ribeiro DA, de Oliveira Monteiro NR,
- 563 Céspedes IC, et al. Physical therapy and psychological intervention normalize cortisol levels
- and improve vitality in women with endometriosis. Journal of Psychosomatic Obstetrics &
- 565 Gynecology 2012;33:191–8. https://doi.org/10.3109/0167482X.2012.729625.
- 566 [73] Gonçalves A, Barros N, Bahamondes L. The Practice of Hatha Yoga for the Treatment
- of Pain Associated with Endometriosis. Journal of Alternative and Complementary Medicine
- 568 (New York, NY) 2016;23. https://doi.org/10.1089/acm.2015.0343.
- 569 [74] Cheong Y, Smotra G, Williams A. Non-surgical interventions for the management of
- 570 chronic pelvic pain. The Cochrane Database of Systematic Reviews 2014;3:CD008797.
- 571 https://doi.org/10.1002/14651858.CD008797.pub2.
- 572 [75] World Health Organization. Programme on Traditional Medicine. WHO traditional
- 573 medicine strategy 2002-2005 2002.
- 574 [76] Burke A, Lam C, Stussman B, Yang H. Prevalence and patterns of use of mantra,
- 575 mindfulness and spiritual meditation among adults in the United States. BMC Complementary
- and Alternative Medicine 2017;17. https://doi.org/10.1186/s12906-017-1827-8.
- 577 [77] Qu S, Olafsrud SM, Meza-Zepeda LA, Saatcioglu F. Rapid Gene Expression Changes
- 578 in Peripheral Blood Lymphocytes upon Practice of a Comprehensive Yoga Program. PLOS
- 579 ONE 2013;8:e61910. https://doi.org/10.1371/journal.pone.0061910.
- 580 [78] Khalsa SB, Hickey Schultz L, Cohen D, Steiner N, Cope S. Evaluation of the Mental
- Health Benefits of Yoga in a Secondary School: A Preliminary Randomized Controlled Trial.
- 582 The Journal of Behavioral Health Services & Research 2011;39:80-90.
- 583 https://doi.org/10.1007/s11414-011-9249-8.

- 584 [79] Sheng JA, Bales NJ, Myers SA, Bautista AI, Roueinfar M, Hale TM, et al. The
- 585 Hypothalamic-Pituitary-Adrenal Axis: Development, Programming Actions of Hormones, and
- 586 Maternal-Fetal Interactions. Frontiers in Behavioral Neuroscience 2021;14.
- 587 https://doi.org/10.3389/fnbeh.2020.601939.
- 588 [80] Granath J, Ingvarsson S, Von Thiele Schwarz U, Lundberg U. Stress Management: A
- Randomized Study of Cognitive Behavioural Therapy and Yoga. Cognitive Behaviour Therapy
- 590 2006;35:3–10. https://doi.org/10.1080/16506070500401292.
- 591 [81] Posadzki P, Ernst E. Yoga for low back pain: a systematic review of randomized clinical
- trials. Clinical Rheumatology 2011;30:1257–62. https://doi.org/10.1007/s10067-011-1764-8.
- 593 [82] Eaves ER, Sherman KJ, Ritenbaugh C, Hsu C, Nichter M, Turner JA, et al. A qualitative
- 594 study of changes in expectations over time among patients with chronic low back pain seeking
- 595 four CAM therapies. BMC Complementary and Alternative Medicine 2015;15:12.
- 596 https://doi.org/10.1186/s12906-015-0531-9.
- 597 [83] Cramer H, Lauche R, Haller H, Dobos G. A Systematic Review and Meta-analysis of
- Yoga for Low Back Pain. The Clinical Journal of Pain 2013;29.
- 599 [84] Jeter PE, Slutsky J, Singh N, Khalsa SBS. Yoga as a Therapeutic Intervention: A
- 600 Bibliometric Analysis of Published Research Studies from 1967 to 2013. The Journal of
- 601 Alternative and Complementary Medicine 2015;21:586–92.
- 602 https://doi.org/10.1089/acm.2015.0057.
- 603 [85] Toussaint L, Nguyen QA, Roettger C, Dixon K, Offenbächer M, Kohls N, et al.
- 604 Effectiveness of Progressive Muscle Relaxation, Deep Breathing, and Guided Imagery in
- 605 Promoting Psychological and Physiological States of Relaxation. Evidence-Based
- 606 Complementary and Alternative Medicine 2021;2021:5924040.
- 607 https://doi.org/10.1155/2021/5924040.
- 608 [86] Ross A, Williams L, Pappas-Sandonas M, Touchton-Leonard K, Fogel D. Incorporating
- Yoga Therapy into Primary Care: The Casey Health Institute. International Journal of Yoga
- 610 Therapy 2015;25:43–9. https://doi.org/10.17761/1531-2054-25.1.43.
- 611 [87] Ranabir S, Reetu K. Stress and hormones. Indian Journal of Endocrinology and
- 612 Metabolism 2011;15.
- 613 [88] Silverman MN, Sternberg EM. Glucocorticoid regulation of inflammation and its
- functional correlates: from HPA axis to glucocorticoid receptor dysfunction. Annals of the New
- 615 York Academy of Sciences 2012;1261:55–63. https://doi.org/10.1111/j.1749-
- 616 6632.2012.06633.x.

617	[89] Kinlein SA, Wilson CD, Karatsoreos IN. Dysregulated Hypothalamic-Pituitary-
618	Adrenal Axis Function Contributes to Altered Endocrine and Neurobehavioral Responses to
619	Acute Stress. Frontiers in Psychiatry 2015;6. https://doi.org/10.3389/fpsyt.2015.00031.
620	[90] Gopal A, Mondal S, Gandhi A, Arora S, Bhattacharjee J. Effect of Integrated Yoga
621	Practice on immune response in examination stress- A preliminary Study. International Journal
622	of Yoga 2011;4:26–32. https://doi.org/10.4103/0973-6131.78178.
623	[91] Sarkar G, Parui SK, Banerjee S. A Dielectric Resonator MIMO Antenna for Intelligent
624	Transportation Systems, 2021, p. 3–9. https://doi.org/10.1007/978-981-15-9433-5_1.
625	[92] Gowda S, .H R, Silvia C. Molecular Signature of the Immune Response to Yoga
626	Therapy in Stress-related Chronic Disease Conditions: An Insight. International Journal of
627	Yoga 2019;13. https://doi.org/10.4103/ijoy.IJOY_82_18.
628	[93] Garavaglia E, Ricci E, Chiaffarino F, Cipriani S, Cioffi R, Viganò P, et al. Leisure and
629	occupational physical activity at different ages and risk of endometriosis. European Journal of
630	Obstetrics & Gynecology and Reproductive Biology 2014;183:104-8.
631	https://doi.org/10.1016/j.ejogrb.2014.10.031.
632	
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644 Table 1: Staging of endometriosis stages, (Zondervan et al. 2022).

Stage	Classification	Description
1 2	Minimal Mild	A small number of superficial implants and mild adhesions More, slightly deeper implants
3	Moderate	Many deep implants, small endometriomas on one or both ovaries, and some fine adhesions
4	Severe	Many deep implants, large endometriomas on one or both ovaries, and many dense adhesions, sometimes with the rectum adhering to the posterior part of the uterus.

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647 GnRH: gonadotropin-realising hormone.

648 BMD: Bone minerl density

649 AP 650 CD:

651 PMK: pacemaker652 PA: physical activity

Table 1 21: Summary of data with physical activity effects on endometriosis.

Studies	Age	Symptom	Diagnostic	particularities	Intervention	Contribution of literature	Level of evidence
		Symptomatic Symptomatic	Laparoscopic diagnosis	Premenopausal / Untreated	GnRH analogue for 6 months	GnRH + brisk walking + aerobic training: femoral neck 6 months: limited loss of BMD At 12 months: increase in BMD	strong evidence (p=0.029)
Bergström et al. [63]	23-28 yrs				aerobic walking and training 1h/week 12 months	GnRH + brisk walking + aerobic training: spine At 6 months: limited loss of BMD At 12 months: BMD increase	weak evidence (p=0.30)
Carpenter	Average 31 yrs	Symptomatic diagnosi	Laparoscopic diagnosis Danazol treatment	Danazol	Aerobic fitness at 50-70% of the maximum heart rate 40 minutes	AP + relaxation: decrease in side effects (for AFS 6 score 3)	strong evidence (p=0,007)
et al. [64]				4 times/week 24 weeks	AP + flexibilities: lower testosterone	weak evidence (P<0.020)	
						Less weight gain	weak evidence (p=0.12)
	Average	Symptomatic	Laparoscopic	none	Halta yoga		strong evidence
Evans et al. [10]	36.8 yrs	(Analog visual	diagnosis		2h	Halta yoga: reduction in pain	(p<0.001)
		scale)			2 times/week 8 weeks		(P 0.001)
	18-48 yrs	not specified	not specified	1 dose of letuprolide	Progressive muscle relaxation (PMR) 40 minutes	PMK: pain improvement, anxiety, depression, fatigue	strong evidence

					2 times/week - 12 weeks Home practice: 2/day (relaxation brochure, CD)		(p<0.001)
			Laparoscopic diagnosis			Professional "standing" PA: protective effect of endometriosis	weak evidence (p=0.22)
	18-76 yrs.					Leisure PA: protective effect of	strong evidence
Garavaglia et al. [93]	Mean = 35 yrs.			not specified	Interview with physical activity intensity and leisure	endometriosis	(p=0.0006)
						Moderate/Intense PA: increases risk of endometriosis	weak evidence (p=0.08)
						IMC ≤ 25 increase women with endometriosis	weak evidence (P=0.07)
Mira et al. [44]	Not specified	Not specified	Not specified	GnRH	Walking or aerobic training 6 months	Increased BMD gain rate in the femur after completion of treatment and prevents bone loss while taking medication	Not specified
	13 to 50 years	Symptomatic	Not specified	Treatment leuprolide	Jacobson method 12 weeks	Improvement of anxiety indices, depression, quality of life	weak evidence (p<0.05)
	Not specified			Danazol	Specific Exercises 24 weeks	Decreased pelvic symptoms in both groups	(No statistical difference)

						Recurrence of endometriosis later (16 months vs. 11 months)	weak evidence (Not significant)
						Decreased testosterone level	strong evidence (p<0.020)
	13 to 50 years		Surgery	Not specified	Halta yoga + postures + relaxation + breathing 2 times/week 8 weeks	Pain relief and quality of life improvement with no change in menstrual flow	Not specified
Ricci et al.[43]	Not specified	Not specified	Not specified	infertility clinic / Not specified	Not specified	PA can reduce the risk of endometriosis	Weak evidence (Not significant)
				Previous	Halta yoga + postures + relaxation + breathing + meditation	Halta yoga + relaxation: reduction in pain	strong evidence $(p < 0.001)$
Tennfjord et al.[8]	34,88 ± 6,70			hormonal and surgical treatment	2 hours 2 times/week 8 weeks	Halta yoga: improvement of control, impotence, well-being, emotional and self-image (EHP-307)	strong evidence (p < 0.001
	32.0 ± 1.3 yrs.	Symptomatic 7 years of delay in taking charge		No effect of medical or surgical treatment	Body awareness, breathing exercise, stretching, general movement, PFM strength + cognitive behavioral therapy	PA + cognitive approach: improvement in the areas of vitality and physical functioning	strong evidence (p<0,05)

		1h + 1h30 1 time/Week 10 weeks	Decrease cortisol level	strong evidence (p=0.04)
Ensari et al.[53] Ensari et al.[53] 1009 participan ts 38 countries, Range 14.3–62.9	70% (N=702) had laparoscopic confirmation of their diagnosis, 19.8% (N=200) had a clinician diagnosis and 10.6% (N=107) had suspected endometriosi s	-Prevalence of self-reported exercise modalities in the study samplecardiovascular'activities -Muscle strength and endurance multimodal exercises (eg, high-intensity interval training of both -cardiovascular and muscular endurance) -fitness activities	-association of previous-day (day <i>t-1</i>) exercise with pain symptoms on day <i>t</i> was moderated by habitual exercise frequency -indicating that those who regularly exercise at least ~3 times per week are less likely to report pain symptoms after having exercised on the previous day.	strong evidence (p<0.0007) for day-level pain score, p=0.026 for Δ-score).