



March Is Endometriosis  
*Awareness*  
Month

ENDOMETRIOSIS  
BULLETIN  
APRIL 2022 /  
ISSUE XXI

[www.endometriozisdernegi.org](http://www.endometriozisdernegi.org)

## SELECTED ARTICLES

New Treatments for  
endometriosis

A new diagnostic test for  
endometriosis: saliva testing

Risk of preterm birth  
in endometriosis

Special  
Interview



Mario Malzoni

### PREFACE

Hello,

We are here with you again with our 21<sup>st</sup> issue. You can find the newest updates on endometriosis and adenomyosis along with our EndoMarch activities here in this bulletin, which is published every three months.

In this issue, you can find updates on the treatment of endometriosis, results of a promising study on the role of saliva analysis in the diagnosis of endometriosis, relationship between endometriosis/adenomyosis and migraine, applications of artificial intelligence on endometriosis and a study on the relationship between endometriosis and preterm labor.

March is endometriosis awareness month. All over the world, various events are held during this month to raise awareness on endometriosis. With the joint effort of The Turkish Endometriosis and Adenomyosis Society's Board Members and Junior Group, we organized various scientific and social events to raise awareness among our patients and the Turkish population. Details regarding our events can be found in this issue. We are proud to be closely followed and appreciated by The Worldwide Endomarch Group.

Since endometriosis is a chronic disease, various studies are carried out all around the world in order to find solutions to ease the daily lives of endometriosis patients. We, as The Turkish Society, wanted to draw attention to the importance of this issue by publishing a public announcement at the beginning of March in order to get endometriosis accepted among the category of chronic diseases in Turkey. We hope that the lives of our endometriosis patients can be made a little easier by taking the necessary steps regarding this issue in near future.

The European Society of Human Reproduction and Embryology updated its endometriosis guideline in 2022 with the contribution of expert scientists. Our association's founding president, Prof. Engin Oral, MD and our treasurer, Assoc. Prof. Pinar Yalcin Bahat, MD, contributed to the medical treatment of endometriosis associated pain section of this guideline. You can visit <https://www.eshre.eu/Guidelines-and-Legal/Guidelines/Endometriosis-guideline.aspx> to access the guideline and patient leaflet.

34<sup>th</sup> and 35<sup>th</sup> live webinar and Instagram series on endometriosis were held with the participation of **Koray Elter, Funda Gode, Seher Sari Kayalarli, Fitnat Topbas Selcuki**.

During this period, January, February and March, the webinar series organized by the European Endometriosis League continued with the valuable presentations of **Eliana Montanari, Laurin Burla** and **Sun-Wei Guo**.

On 20-21 January 2022 in Dubai, our president **Taner Usta** and our board members **Engin Oral** and **Ahmet Kale** contributed to the 7<sup>th</sup> EMEL Conference on Endometriosis and Uterine Disorders as speakers and with a live endometriosis surgery broadcast from Istanbul.

Our new webinar series, **Experts Discuss Endo-Adeno with Real Cases**, moderated by **Engin Oral** will be held on April 5<sup>th</sup> with the participation of **Yucel Karaman, Umit Inceboz, Bulent Berker, Yusuf Aytac Tohma**.

Among the upcoming meetings of our society, 3<sup>rd</sup> **International Endometriosis School Istanbul** will be chaired by **Engin Oral, Taner Usta** and **Ertan Saridogan**, on **May 27-28**, with the participation of many experts in the field of endometriosis from Turkey and abroad. In addition to the theoretical training, the participants will be given hands-on laparoscopic surgery training in the animal laboratory. We will hold the 14<sup>th</sup> of our EndoAcademy meetings in Adana in June. The meeting, which will be chaired by **Turan Cetin** and **Cihan Kaya**, will cover the diagnosis of endometriomas, its medical and surgical treatment, its relationship with infertility and pelvic pain.

Endo-Specialist interview guest of this month's issue was **Mario Malzoni** from Italy. **Ezgi Darici**, one of the Junior Group members, conducted this valuable interview with him.

We hope to be together again with developments from the world of endometriosis and adenomyosis in our next issue.

Best Regards,  
Prof. Taner Usta, MD  
President of the Endometriosis & Adenomyosis Society

## Turkish Endometriosis & Adenomyosis Society Board of Directors 2019-2022

Turkish Endometriosis and Adenomyosis Society's Founding President  
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**Kaya, MD**

Endometriosis e-bulletin is prepared by Turkish Endometriosis & Adenomyosis Society. If there are any topics that you would like us to include in the bulletin or any questions you would like to ask, you can contact us via e-mail at [drcihankaya@gmail.com](mailto:drcihankaya@gmail.com).

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## PUBLIC ANNOUNCEMENT ON WHY ENDOMETRIOSIS SHOULD BE CONSIDERED AMONG THE CHRONIC DISEASES



### Announcement about the acceptance of endometriosis (chocolate cyst) as a chronic disease:

Chocolate cyst disease (**endometriosis**) is a chronic, estrogen dependent, progressive disease defined as the presence of cells belonging to the inner lining of the uterus, **outside of it**. The disease commonly affects ovaries, tubes, ligaments fixating the uterus, and rarely intestines, bladder, urethra, nerves, and old surgical incisions.

Although it can be seen at any age, in our country, approximately **2 million of 20 million** women of reproductive age are at risk for this disease. The diagnosis of the disease takes an **average of 7 years**, even in developed countries. It can cause pain often **during menstruation, sexual intercourse, defecation and urination, also may lead to infertility**. These symptoms of the disease affect women's quality of life, their place in the work force, and success of schoolgirls negatively. Since it is a **progressive disease**, it may lead to **loss of organs** such as ovaries, uterus, intestines, bladder and kidneys, and may also **affect pregnancy and fertility**. The disease's need for **continuous treatment, its chronic course, its recurrence despite treatment, although it may seem like a benign disease, its spread to the surrounding organs, the risk of cancer development, the need for more than one surgery in some patients and the chronic pain causes loss of work force and deterioration in quality of life**. It isolates women from social life, especially at the age when they should be productive for the society, disrupts wellbeing of the family. Thus, it creates a **great social-economic loss** for our country.

The above-mentioned features of the disease reveal the necessity of evaluating endometriosis in the category of diseases such as **diabetes, asthma, COPD, hypertension and even cancer**.

If we consider the importance of women in family and business life in Turkey, we, as the Society of **Endometriosis and Adenomyosis**, request on behalf of the public that this disease, which is **common and causes organ loss** due to neglect, be included in the category of diseases requiring chronic care and health authorities to support studies on endometriosis.

**Society of Endometriosis and Adenomyosis**

## TABLE OF CONTENTS

### **A** SELECTED ARTICLES

**1. New Therapeutics in Endometriosis: A Review of Hormonal, Non- Hormonal, and Non-Coding RNA Treatments**

Brichant G, Laraki I, Henry L, Munaut C, Nisolle M. *Int J Mol Sci.* 2021;22(19):10498.

**2. Machine learning algorithms as new screening approach for patients with endometriosis**

Bendifallah S, Puchar A, Suisse S, et al. *Sci Rep.* 2022;12(1):639.

**3. Mapping of endometriosis in patients with unilateral endometrioma**

RSDC, Maia SB, Lúcio JD, Lima MD, Ribeiro HSAA, Ribeiro PAAG. *Medicine (Baltimore).* 2021;100(33):e26979.

**4. Salivary MicroRNA Signature for Diagnosis of Endometriosis**

Bendifallah S, Suisse S, Puchar A, et al. *J Clin Med.* 2022;11(3):612.

**5. Migraine Is More Prevalent in Advanced-Stage Endometriosis, Especially When Co-Occurring with Adenomyosis**

Wu Y, Wang H, Chen S, et al. *Front Endocrinol (Lausanne).* 2022;12:814474.

**6. Endometriosis and preterm birth: A Danish cohort study**

Breintoft K, Arendt LH, Uldbjerg N, Glavind MT, Forman A, Henriksen TB. *Acta Obstet Gynecol Scand.* 2022;00:1–7.

### **B- NEWS FROM OUR SOCIETY**

### **C- NEWS FROM THE WORLD OF ENDOMETRIOSIS**

### **D- INTERVIEW WITH AN ENDO SPECIALIST**

### **E- ARTICLES ON ENDOMETRIOSIS FROM OUR COUNTRY FROM THE LAST THREE MONTHS**

### **F- SOCIAL MEDIA**

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## A SELECTED ARTICLES

### 1 New Therapeutics in Endometriosis: A Review of Hormonal, Non-Hormonal, and Non-Coding RNA Treatments

Brichant G, Laraki I, Henry L, Munaut C, Nisolle M. *Int J Mol Sci.* 2021;22(19):10498.

#### Abstract

Endometriosis is defined as endometrial-like tissue outside the uterine cavity. It is a chronic inflammatory estrogen-dependent disease causing pain and infertility in about 10% of women of reproductive age. Treatment nowadays consists of medical and surgical therapies. Medical treatments are based on painkillers and hormonal treatments. To date, none of the medical treatments have been able to cure the disease and symptoms recur as soon as the medication is stopped. The development of new biomedical targets, aiming at the cellular and molecular mechanisms responsible for endometriosis, is needed. This article summarizes the most recent medications under investigation in endometriosis treatment with an emphasis on non-coding RNAs that are emerging as key players in several human diseases, including cancer and endometriosis.



**Keywords:** endometriosis, non-coding RNA, cell migration, cell proliferation, apoptosis, fibrosis, angiogenesis, stem cells, inflammation

### 2 Machine learning algorithms as new screening approach for patients with endometriosis

Bendifallah S, Puchar A, Suisse S, et al. *Sci Rep.* 2022;12(1):639.

#### Abstract

Endometriosis—a systemic and chronic condition occurring in women of childbearing age—is a highly enigmatic disease with unresolved questions. While multiple biomarkers, genomic analysis, questionnaires, and imaging techniques have been advocated as screening and triage tests for endometriosis to replace diagnostic laparoscopy, none have been implemented routinely in clinical practice. We investigated the use of machine learning algorithms (MLA) in the diagnosis and screening of endometriosis based on 16 key clinical and patient-based symptom features. The sensitivity, specificity, F1-score and AUCs of the MLA to diagnose endometriosis in the training and validation sets varied from 0.82 to 1, 0-0.8, 0-0.88, 0.5-0.89, and from 0.91 to 0.95, 0.66-0.92, 0.77-0.92, respectively. Our data suggest that MLA could be a promising screening test for general practitioners, gynecologists, and other front-line health care providers. Introducing MLA in this setting represents a paradigm change in clinical practice as it could replace diagnostic laparoscopy. Furthermore, this patient-based screening tool empowers patients with endometriosis to self-identify potential symptoms and initiate dialogue with physicians about diagnosis and treatment, and hence contribute to shared decision making.



### 3 Mapping of endometriosis in patients with unilateral endometrioma

Araujo RSDC, Maia SB, Lúcio JD, Lima MD, Ribeiro HSAA, Ribeiro PAAG. *Medicine (Baltimore)*. 2021;100(33):e26979.

#### Abstract

To map the distribution of the sites most affected by endometriosis in patients with unilateral ovarian endometriomas. A descriptive case series of 84 patients with unilateral endometriomas undergoing laparoscopy for the treatment of endometriosis. To evaluate the distribution of the sites of endometriosis lesions, the peritoneal compartments were divided into 5 zones: zone 1/the anterior compartment, including the anterior uterine serosa, vesicouterine fold, round ligament, and bladder; zone 2/the lateral compartment, including the left and right ovary, ovarian fossa, tubes, mesosalpinx, uterosacral ligaments, parametrium, and the ureter; zone 3/the posterior compartment, including posterior uterine serosa, the pouch of Douglas, posterior vaginal fornix, and bowel; zone 4 consisting of the abdominal wall; and zone 5 consisting of the diaphragm. Of the 5 zones evaluated, the lateral compartment (zone 2) was the most affected, with 60.7% of the patients having dense adhesions around the left ovarian fossa and 57.1% around the right ovarian fossa. The ovarian endometriomas were more commonly found



on the left side (54.8%) compared to the right (45.2%). In the posterior compartment (zone 3), the posterior cul-de-sac was obliterated in 51.2% of the patients. In the anterior compartment (zone 1), there were lesions in the vesicouterine fold in 30.9% of the patients and in the bladder in 19%. Lesions were found in the abdominal wall (zone 4) and diaphragm (zone 5) in 21.4% and 10.7% of patients, respectively. Unilateral endometriomas are important markers of the severity of endometriosis.

### 4 Salivary MicroRNA Signature for Diagnosis of Endometriosis

Bendifallah S, Suisse S, Puchar A, et al. *J Clin Med*. 2022;11(3):612.

#### Abstract

**Background:** Endometriosis diagnosis constitutes a considerable economic burden for the healthcare system with diagnostic tools often inconclusive with insufficient accuracy. We sought to analyze the human miRNAome to define a saliva-based diagnostic miRNA signature for endometriosis.

**Methods:** We performed a prospective ENDO-miRNA study involving 200 saliva samples obtained from 200 women with chronic pelvic pain suggestive of endometriosis collected between January and June 2021. The study consisted of two parts: (i) identification of a biomarker based on genome-wide miRNA expression profiling by small RNA sequencing using next-generation sequencing (NGS) and (ii) development of a saliva-based miRNA diagnostic signature according to expression and accuracy profiling using a Random Forest algorithm.

**Results:** Among the 200 patients, 76.5% (n = 153) were diagnosed with endometriosis and 23.5% (n = 47) without (controls). Small RNA-seq of 200 saliva samples yielded ~4642 M raw sequencing reads (from ~13.7 M to ~39.3 M reads/sample). Quantification of the filtered reads and identification of known miRNAs yielded ~190 M sequences that were mapped to 2561 known miRNAs. Of the 2561 known miRNAs, the feature selection with Random



Forest algorithm generated after internally cross validation a saliva signature of endometriosis composed of 109 miRNAs. The respective sensitivity, specificity, and AUC for the diagnostic miRNA signature were 96.7%, 100%, and 98.3%.

**Conclusions:** The ENDO-miRNA study is the first prospective study to report a saliva-based diagnostic miRNA signature for endometriosis. This could contribute to improving early diagnosis by means of a non-invasive tool easily available in any healthcare system.

**Keywords:** diagnostic; endometriosis; miRNA; saliva; signature.

## 5 Migraine Is More Prevalent in Advanced-Stage Endometriosis, Especially When Co-Occurring with Adenomyosis

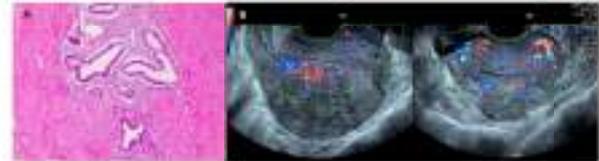
Wu Y, Wang H, Chen S, et al. *Front Endocrinol (Lausanne)*. 2022;12:814474.

### Abstract

**Background:** Emerging data suggest a significant association between migraine and endometriosis, however the relationship between migraine and endometriosis severity or adenomyosis is unclear. Our objectives were to explore the relationship between migraine and endometriosis, according to the endometriosis severity and co-exist with adenomyosis or not.

**Methods:** This case-control study of 167 endometriosis patients verified by surgery and 190 patients for other benign gynecological conditions (control subjects) was performed from September 2017 and January 2021. There is 49 adenomyosis detected by transvaginal ultrasound or histologic diagnosis among the endometriosis patients. Besides, we also included 41 adenomyosis but without endometriosis patients as a subgroup. All women completed a self-administered headache questionnaire and diagnosed as migraine according to the International Headache Society classification. The severity and stage of endometriosis was evaluated with revised American Society of Reproductive Medicine (rASRM) score. We used logistic regression to estimate the association between the presence of migraine and endometriosis severity while accounting for important confounders, including age, body mass index (BMI) and family history of migraine. We also estimate the risk of adenomyosis alone and adenomyosis with co-occurring endometriosis in migrainous women.

**Results:** Migraine was significantly more prevalent in endometriosis patients compared with controls (29.9% vs. 12.1%,  $p < 0.05$ ), but the prevalence was similar between isolated



adenomyosis patients and controls (9.8% vs. 12.1%,  $p > 0.05$ ). For all endometriosis and control participants, migraineurs were 4.6-times (OR=4.6; 95% CI 2.7-8.1) more likely to have severe endometriosis. However, the strength of the association decreased when the analysis examined in moderate stage (OR=3.6, 95% CI 2.1-6.2). The risk of mild and minimal endometriosis was not significant (OR=1.9, 95% CI 0.9-4.0; OR=1.6, 95% CI 0.8-3.4; respectively). When we divided the endometriosis patients according to whether co-occurring with adenomyosis. We found in migrainous women, the risk of endometriosis co-exist with adenomyosis increased, with nearly fivefold greater odds compared with control (OR=5.4; 95% CI 3.0-9.5), and nearly two times higher than the risk of endometriosis without co-exist adenomyosis patients (OR=2.2; 95% CI 1.2-3.8).

**Conclusion:** Our study supports the strong association between migraine and endometriosis. We found migrainous women suffer more frequently from severe endometriosis, especially endometriosis with co-occurring adenomyosis. It is advisable to heighten suspicion for patients who presenting with either these conditions in order to optimize therapy.

**Keywords:** adenomyosis; endometriosis (EM); endometriosis severity; migraine; rASRM score.

## 6 Endometriosis and preterm birth: A Danish cohort study

Breintoft K, Arendt LH, Uldbjerg N, Glavind MT, Forman A, Henriksen TB. *Acta Obstet Gynecol Scand*. 2022;00:1-7.

### Abstract

**Introduction:** Emerging evidence shows that women with endometriosis face a higher risk of preterm birth. However, the pathways are unclear. The objective of this study is to further investigate at different gestational ages the association between endometriosis and different pathways of preterm birth including, medically indicated preterm birth, premature pre-labor rupture of membranes (PPROM), and spontaneous labor contractions.

**Material and methods:** In this population-based cohort study we linked singleton pregnancies from the Aarhus Birth Cohort to the Danish National Patient Registry, the Danish Medical Birth Registry, the Danish National Pathology Registry and Data Bank, and the Danish in vitro fertilization registry to gather information on endometriosis status, outcomes and maternal characteristics. We investigated preterm birth before 37 completed weeks of gestation and very preterm birth before 32 completed weeks of gestation. We explored different pathways including medically indicated preterm birth defined as induction of labor with intact



membranes and no prior labor contractions, PPRM defined as rupture of membranes, and spontaneous labor contractions defined as contractions with intact membranes resulting in labor.

**Results:** We found that women with endometriosis had an increased risk of preterm birth before 37 gestational weeks overall (adjusted hazard rate [aHR] 1.6, 95% confidence interval [CI] 1.3-1.9) and very preterm birth before 32 gestational weeks (aHR 1.8, 95% CI 1.1-2.9) compared with women without

endometriosis. Medically indicated preterm birth was more prominent in women with endometriosis in deliveries before 37 gestational weeks (aHR 2.4, 95% CI 1.8-3.2) whereas spontaneous labor contractions were more common before 32 gestational weeks (aHR 2.2, 95% CI 1.1-4.5) in women with endometriosis compared with women without endometriosis. Further, in the analyses restricted to women with a histologically verified diagnosis of endometriosis, the results were strengthened overall and showed that women with endometriosis had an increased risk of PPROM before 32 gestational weeks (aHR 3.49, 95% CI 1.36-8.98).

**Conclusions:** Endometriosis was associated with both preterm and very preterm birth; however, apparently through different pathways. Women with endometriosis were more prone to have medically indicated preterm births before 37 gestational weeks and spontaneous preterm births before 32 gestational weeks compared with women without endometriosis.

**Keywords:** endometriosis; induced labor onset premature birth preterm premature rupture of the membranes; labor.

## B NEWS FROM OUR SOCIETY PAST ACTIVITIES

### @endometriosis\_tr Live Broadcasts

On our society's Instagram account, we continued our live broadcasts, which are taking place since the beginning of the pandemic. During the last three months, we went live with our 34<sup>th</sup> and 35<sup>th</sup> broadcasts with the participation of Prof. Koray Elter, MD, Assoc. Prof. Funda Gode, MD, Seher Sari Kayalarli, MD, and Fitnat Topbas Selcuki, MD.

**Soru-Cevap 34**  
Endometriosis hakkında merak edilenleri cevaplayacağız  
Tarih: 19.01.2022 Çarşamba  
Saat: 21:00  
Konum: @endometriosis\_tr Instagram Hesabı

**LIVE** **LIVE**

Prof. Dr. Koray Elter Dr. Seher Sari Kayalarli

**Q&A 34**  
**Everything You Want to Know About Endometriosis**  
Prof. Koray Elter, MD  
Seher Sari Kayalarli, MD

**Soru-Cevap 35**  
Endometriosis hakkında merak edilenleri cevaplayacağız  
Tarih: 09.02.2022 Çarşamba  
Saat: 21:00  
Konum: @endometriosis\_tr Instagram Hesabı

**LIVE** **LIVE**

Dr. Dr. Funda Gode Dr. Fitnat Topbas Selcuki

**Q&A 35**  
**Everything You Want to Know About Endometriosis**  
Prof. Funda Gode, MD  
Fitnat Topbas Selcuki, MD

The second of our webinar series 'Experts Discuss Endo-Adeno with Real Cases' will take place on the 5<sup>th</sup> of April, with Prof. Yucel Karaman, MD, Prof. Umit Inceboz, MD, Prof. Bulent Berker, MD participating and Prof. Engin Oral, MD moderating.

**GERÇEK OLGULARLA ENDO&ADENO UZMANLAR TARTIŞIYOR**

**5 NİSAN 2022 SALI 21:00 - 22:30**

**KAYIT OLMAK İÇİN** [www.fisakademi.org](http://www.fisakademi.org)

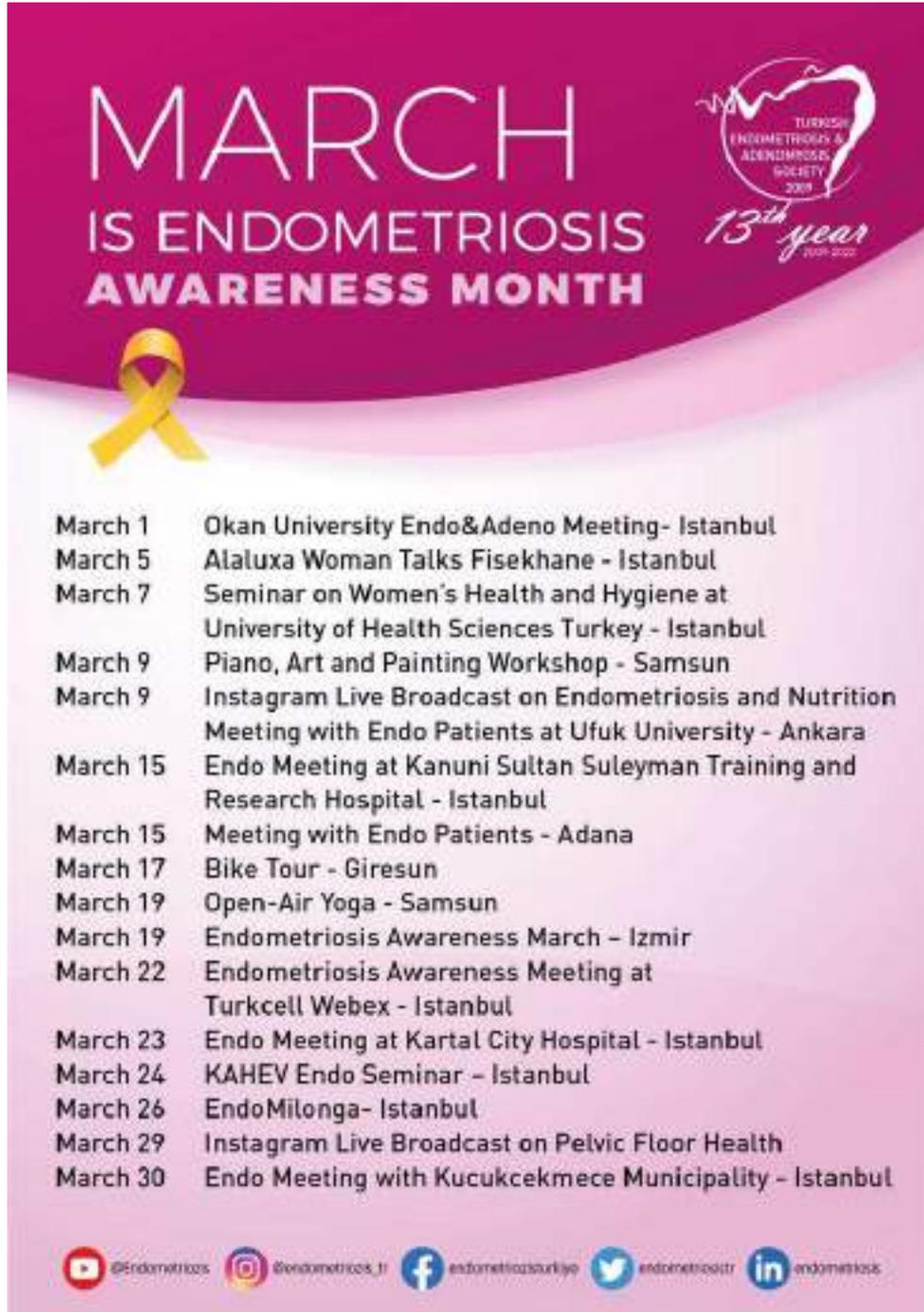
Prof. Dr. Yucel Karaman Prof. Dr. Umit Inceboz Prof. Dr. Bulent Berker Doç. Dr. Yavuz Altac Tokmak

Prof. Dr. Engin Oral

[www.info@endometriozisdernegi.org](http://www.info@endometriozisdernegi.org) | OLGULAR: EndoGenç Grup / Uzm. Dr. Ayşegül BESTEL

### EndoMarch Activities

March is endometriosis awareness month. During this month we organized both educational and social activities through the country. We held seminars on endometriosis awareness at several universities and dormitories. Additionally, we organized a march for endometriosis, several workshops, live Q&As on our Instagram account and prepared endometriosis awareness billboards which were advertised throughout some of the major cities. All the activity links and videos can be found on our website.



The poster features a pink and white background with a yellow ribbon symbol. The main text reads "MARCH IS ENDOMETRIOSIS AWARENESS MONTH". In the top right corner, there is a logo for the "TURKISH ENDOMETRIOSIS & ADENOMIOSIS SOCIETY" established in 2009, celebrating its 13th year (2009-2022). Below the main text, a list of activities is provided for each day of the month. At the bottom, there are social media icons for YouTube, Instagram, Facebook, Twitter, and LinkedIn, each with its respective handle.

**MARCH IS ENDOMETRIOSIS AWARENESS MONTH**

TURKISH ENDOMETRIOSIS & ADENOMIOSIS SOCIETY 2009  
13<sup>th</sup> year 2009-2022

**March 1** Okan University Endo&Adeno Meeting- Istanbul  
**March 5** Alaluxa Woman Talks Fisekhane - Istanbul  
**March 7** Seminar on Women's Health and Hygiene at University of Health Sciences Turkey - Istanbul  
**March 9** Piano, Art and Painting Workshop - Samsun  
**March 9** Instagram Live Broadcast on Endometriosis and Nutrition Meeting with Endo Patients at Ufuk University - Ankara  
**March 15** Endo Meeting at Kanuni Sultan Suleyman Training and Research Hospital - Istanbul  
**March 15** Meeting with Endo Patients - Adana  
**March 17** Bike Tour - Giresun  
**March 19** Open-Air Yoga - Samsun  
**March 19** Endometriosis Awareness March - Izmir  
**March 22** Endometriosis Awareness Meeting at Turkcell Webex - Istanbul  
**March 23** Endo Meeting at Kartal City Hospital - Istanbul  
**March 24** KAHEV Endo Seminar - Istanbul  
**March 26** EndoMilonga- Istanbul  
**March 29** Instagram Live Broadcast on Pelvic Floor Health  
**March 30** Endo Meeting with Kucukcekmece Municipality - Istanbul

 @Endometriosis  @endometriosis\_tr  endometriosisurkiye  endometriocentr  endometriosis













 ENDOMETRİOZİS VE  
ADENOMYOZİS DERNEĞİ  
YÖNETİM KURULU

**TÜM SORULARINIZI  
CEVAPLIYOR!**

**27 MART 2022  
20:00  
ZOOM TOPLANTISI**

info@endometriosisdernegi.org

**İLETİŞİM** 0532 515 69 99

**MART ENDOMETRİOZİS FARKINDOLİK AYIDIR**



 Prof. Dr. Taner Usta	 Prof. Dr. Ümit İnceboz	 Prof. Dr. Ahmet Kale	
 Prof. Dr. Engin Oval	 Doç. Dr. Pınar Yalçın Rahat	 Doç. Dr. Hale Göksever Çelik	 Doç. Dr. Cihan Kaya

## PLANNED ACTIVITIES

### XIV. EndoAcademy, Endometrioma, Adana

14<sup>th</sup> EndoAcademy-Endometrioma meeting will be held on the 12<sup>th</sup> of June in Adana. The meeting will be chaired by **Prof. Turan Cetin, MD** and **Assoc. Prof. Cihan Kaya, MD**. The meeting will cover diagnosis, medical and surgical treatments, infertility and pelvic pain with a focus on endometrioma. Many experts in the field will share their knowledge and experiences on these topics.







## 3<sup>rd</sup> International **ENDOMETRIOSIS SCHOOL** Istanbul

May 27-28, 2022  
Medtronic Innovation Center  
Istanbul - Turkey

Dear colleagues,

With many countries lifting the COVID-19 pandemic restrictions, we would like to share our happiness with you that the 3rd International Endometriosis School Istanbul will be held in person in Istanbul. After the first two International Endometriosis School meetings, which were successfully held in Istanbul in 2018 and 2019, the Turkish Endometriosis and Adenomyosis Society will organize the 3rd International Endometriosis School meeting under the chairmanship of Dr. Taner Usta, Dr. Engin Oral, and Dr. Ertan Sarıdoğan.

The meeting, which aim to attract the attention of young specialists and researchers especially interested in endometriosis, consists of four main topics. All lectures will be given by well-known endometriosis experts. The first part will cover the classification, imaging, and diagnosis of endometriosis. The second part will consist of the topics related to infertility, pregnancy outcomes, and pain management in endometriosis patients. The third part will include the surgical management of endometriosis in a broad aspect, and finally, the fourth part will cover recent guidelines, management of recurrent cases, and surgical complications. The attendees will also have the opportunity of watching a live deep infiltrating endometriosis surgery.

Additionally, hands-on practice session held in the animal laboratory under the supervision of expert trainers will aim to improve participants' surgical skills. Exercises will include basic endoscopic training, hysterectomy and variety of advanced techniques related to endometriosis, such as bowel resection and anastomosis, ureteroneocystostomy, and management of major vascular injuries.

We cordially invite you to join us in this meeting, where we are certain that you will be exposed to new knowledge on endometriosis. We look forward to seeing you in Istanbul.

Kind regards

**Prof. Taner Usta, MD.**  
Course Director

**Prof. Engin Oral, MD.**  
Course Director

**Prof. Ertan Sarıdoğan, MD.**  
Course Director

[www.endometriosis-turkey.org](http://www.endometriosis-turkey.org)

[www.endometriosis-school.com](http://www.endometriosis-school.com)



# 3<sup>rd</sup> International ENDOMETRIOSIS SCHOOL Istanbul

May 27-28, 2022  
Medtronic Innovation Center  
Istanbul - Turkey

**3rd INTERNATIONAL ENDOMETRIOSIS SCHOOL - ISTANBUL**  
Education Program of Endometriosis and Adenomyosis Society of Turkey

**Course Directors:** Taner Usta, Engin Oral, Ertan Sarıdoğan  
**Scientific Secretary:** Tolga Karacan

DAY 1	MAY 27, 2022
08:00-08:20	Registration
08:20-08:30	Opening: The aim and objectives of the International Endometriosis School, <b>Ertan Sarıdoğan</b> Preschool Evaluation of The Participants (Keypad), <b>İşıl Ayhan</b>
<b>SESSION I</b>	<b>CLASSIFICATION, IMAGING AND DIAGNOSIS</b>
Chairs	<b>Levent Şentürk, Pınar Yalçın Bahat</b>
08:30-08:50	Useful Anatomy for Pelvic Laparoscopic Surgery <b>Ahmet Kale</b>
08:50-09:10	Imaging in Endometriosis <b>Sebastian Schaefer</b>
09:10-09:30	Current Approach to Management of Adenomyosis <b>Joerg Keckstein</b>
09:30-09:50	Classification and Staging of Peritoneal Disease, Endometrioma and Deep Infiltrating Endometriosis <b>Ghassan Lotfi</b>
09:50-10:10	Discussion
<b>10:10-10:30</b>	<b>Coffee Break</b>
<b>SESSION II</b>	<b>INFERTILITY, PREGNANCY, PAIN AND ENDOMETRIOSIS AT THE EXTREMES AGES</b>
Chairs	<b>Koray Elter, Nilüfer Akgün</b>
10:30-10:50	Management of Endometriosis and Infertility <b>Engin Oral</b>
10:50-11:10	Obstetric Outcomes in Endometriosis and Adenomyosis <b>Umberto Leone Roberti Maggiore</b>
11:10-11:30	Pain Management in Endometriosis <b>Moamar Al-Jefout</b>
11:30-11:50	Management of Endometriosis at the Extremes of Reproductive Years <b>Ümit İnceboz</b>
11:50-12:00	Discussion

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**3<sup>rd</sup> International  
ENDOMETRIOSIS  
SCHOOL Istanbul**

May 27-28, 2022  
Medtronic Innovation Center  
Istanbul - Turkey

<b>LIVE-1</b>	<b>LIVE TRANSMISSION SESSION</b>
Chairs	Hüsnü Görgeç, Salih Yılmaz
12:00-14:20	Live Surgery (from Istanbul, Turkey) Surgeon, <b>Taner Usta</b>
<b>12:00-13:00</b>	<b>Lunch Break</b>
<b>SESSION III</b>	<b>SURGICAL TECHNIQUES</b>
Chairs	Hüseyin Cengiz, Cihan Kaya
14:20-14:40	Diagnosis of Adenomyosis: Clinical and Imaging <b>Yutaka Osuga</b>
14:40-15:00	Urinary Tract Endometriosis <b>Horace Roman</b>
15:00-15:20	Bowel Endometriosis: Indication and Surgical Techniques <b>Joerg Keckstein</b>
15:20-15:40	Surgical Techniques for Endometriomas <b>Taner Usta</b>
15:40-16:00	Discussion
<b>16:00-16:20</b>	<b>Coffee Break</b>
<b>SESSION IV</b>	<b>GUIDELINES, RECURRENCE AND COMPLICATIONS</b>
Chairs	Murat Sönmezer, Hale Göksever Çelik
16:20-16:40	How Can We Prevent Complications During Endometriosis Surgery: Tips and Tricks <b>Muhammed Mabrouk</b>
16:40-17:00	Management of Recurrent Endometriosis After Surgical Treatment <b>Helder Ferreira</b>
17:00-17:20	ESHRE Guideline 2022 on Endometriosis, What is New <b>Ertan Sarıdoğan</b>
17:20-17:30	Discussion
17:30-17:40	Postschool Evaluation of the Course by the Participants (Keypad) <b>Cihan Kaya</b>

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# 3<sup>rd</sup> International ENDOMETRIOSIS SCHOOL Istanbul

May 27-28, 2022  
Medtronic Innovation Center  
Istanbul - Turkey

DAY 2	MAY 28, 2022
08:30-09:00	Pig Model for Laparoscopy - Evaluation Medtronic
<b>09:00-11:00</b>	<b>HANDS ON TRAINING</b>
<b>Coordinators</b>	<b>Ahmet Kale, Cihan Kaya</b>
<b>Trainers</b>	<b>Ertan Sandogan, Hüsnü Görgeç, Ahmet Kale, Taner Usta, Cihan Kaya, Tolga Karacan</b>
<b>Training on Live Animal Tissue</b> Practice of Different Energy Types in Live Animal Model Bladder Injury and Repair Dissection of the Ureters and Major Pelvic Vessels	
<b>LIVE-2</b>	<b>LIVE TRANSMISSION SESSION</b>
<b>Chairs</b>	<b>Suat Dede, Tolga Karacan</b>
11:00-13:00	Live Surgery (from London, U.K.) Surgeon, <b>Shaheen Khazali</b>
<b>13:00-17:00</b>	<b>HANDS ON TRAINING</b>
<b>Training on Live Animal Tissue</b> Ureteric Injury and Reanastomosis Hysterectomy	
17:00-17:20	Wrap-up <b>Ertan Sandogan, Hüsnü Görgeç, Ahmet Kale, Taner Usta, Cihan Kaya, Tolga Karacan</b>
17:20-17:30	End of the Course - Delivery of Certificates of Attendance

### Registration Fees

Hands-on Training Registration Fee (2 days)	300 Euro
Theoric Sessions Online Registration Fee (1 day)	50 Euro

*Hands-on Training Registration Fee includes theoretic sessions on May 27, 2022 and hands-on training on May 28, 2022. Theoric Sessions Online Registration Fee includes only theoretic sessions on May 27, 2022.*

*For registration or any question please send an e-mail to [endometriosis@fiskonre.org](mailto:endometriosis@fiskonre.org) or contact from this number +90 530 406 60 10.*

*Executive Secretary of Society: Ekin Ezgi Erdoğan, [info@endometriosisderneqi.org](mailto:info@endometriosisderneqi.org) / +90 532 515 69 99*

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## C NEWS FROM THE WORLD OF ENDOMETRIOSIS

### EEL WEBINAR Program 2022

European Endometriosis League (EEL) are continuing with their Webinars in 2022.

**Eliana Montanari, MD, PhD** explained the presurgical use of ultrasonography in diagnosis in the webinar held in January with the title '**Presurgical Ultrasound Prediction of Deep and Ovarian Endometriosis**'. The EEL webinar '**The Role of MR Imaging in the Diagnosis of Deep Endometriosis**', which was held in February, was presented by **Laurin Burla, MD**. In March, **Sun-Wei Guo, MD, PhD** talked about '**Adenomyosis from Pathogenesis to Treatment**'.

**EEL WEBINARS**

Eliana Montanari, MD., PhD.

**PRESURGICAL ULTRASOUND PREDICTION OF DEEP AND OVARIAN ENDOMETRIOSIS**

Moderator: Prof. Dr. Dr. h.c. mult. Hans-Rudolf Tinnberg

DATE: 11 JANUARY 2022  
TIME: 7:00 PM CET

live.euroendometriosis.com

**EEL WEBINARS**

Laurin Burla, MD.

**THE ROLE OF MR IMAGING IN THE DIAGNOSIS OF DEEP ENDOMETRIOSIS**

Moderator: Joseph Nassif, MD, PhD

DATE: 8 FEB 2022  
TIME: 7:00 PM CET

live.euroendometriosis.com

**EEL WEBINARS**

Sun-Wei Guo, MD., PhD.

**ADENOMYOSIS FROM PATHOGENESIS TO TREATMENT**

Moderator: Stefano Angioni, MD, PhD

DATE: 8 MARCH 2022  
TIME: 7:00 PM CET

live.euroendometriosis.com

European Endometriosis League | GEDÉON RICHTER 120 years in Germany

EEL Webinar series will continue in 2022. For more information, visit <https://www.endometriosis-league.eu/home> or follow the European Endometriosis League or Euro Endo League accounts on social media.

ESHRE 2022 the new Endometriosis Guideline has been published.  
<https://www.eshre.ed/Guidelines-and-Legal/Guidelines/Endometriosis-guideline.aspx>.

European Society of Human Reproduction and Embryology

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**Endometriosis**  
Guideline of European Society of Human Reproduction and Embryology

2022  
ESHRE Endometriosis Guideline Development Group



## Table of Contents

Disclaimer	2
<b>Table of Contents</b>	<b>3</b>
<b>Introduction</b>	<b>6</b>
Clinical need	6
Previous versions and significant changes	6
Target users of the guideline	7
Guideline scope	7
Patient population	8
Terminology and definitions	8
<b>List of all recommendations</b>	<b>9</b>
<b>List of research recommendations</b>	<b>18</b>
<b>I. Diagnosis of endometriosis</b>	<b>20</b>
I.1. Signs and symptoms	21
I.2. Diagnostic work-up	24
I.2.a. Clinical examination	24
I.2.b. Medical technologies	25
I.2.c. Diagnostic laparoscopy or empirical treatment	29
I.3. Long term monitoring	31
I.4. Impact of the time of diagnosis on quality of life	34
<b>II. Treatment of endometriosis-associated pain</b>	<b>35</b>
II.1. Analgesics	35
II.2. Hormone treatments	37
II.2.a. Combined hormonal contraceptives	38
II.2.b. Progestagens (including progestogen-only contraceptives) and anti-progestagens	39
II.2.c. GnRH agonists	42
II.2.d. GnRH antagonists	43
II.2.e. Aromatase inhibitors	44
II.3. Surgical treatment	48
II.3.a. Surgery versus diagnostic laparoscopy/medical treatment	48
II.3.b. Ablation versus excision of endometriosis	50
II.3.c. Superficial peritoneal endometriosis	50
II.3.d. Surgical interruption of pelvic nerve pathways	51
II.3.e. Surgery for ovarian endometrioma	51
II.3.f. Surgery for deep endometriosis	53
II.3.g. Nerve-sparing laparoscopy	60
II.3.h. Hysterectomy for endometriosis-associated pain	60
II.3.i. Patient selection for surgery	61
II.4. Medical therapies adjunct to surgery	67
II.4.a. Preoperative medical treatment	67
II.4.b. Postoperative medical treatment	68
II.5. Medical versus surgical treatment for endometriosis	69
II.6. Non-medical management strategies	71
II.6.a. Acupuncture	72
II.6.b. Physical therapies	75
II.6.c. Electrotherapy	75
II.6.d. Psychological interventions	75
II.6.e. Nutrition	77
II.6.f. Traditional Chinese Medicine	78
<b>III. Treatment of endometriosis-associated infertility</b>	<b>81</b>
III.1. Medical treatment	81
III.1.a. Ovarian suppression	81
III.1.b. Hormone or medical therapies as an adjunct to surgical therapy	82
Guideline Endometriosis - 2022	3



III.1.c. Other medical treatments	83
III.2. Surgical treatment	85
III.2.a. Peritoneal endometriosis	85
III.2.b. Ovarian endometriosis	85
III.2.c. Deep endometriosis	86
III.3 Assessing the need for assisted reproduction after surgery	89
III.4. Medically assisted reproduction	90
III.4.a. Intrauterine insemination in women with endometriosis	90
III.4.b. Assisted reproductive technology in women with endometriosis	91
III.5. Medical therapies as an adjunct to MAR	96
III.6. Surgical therapies as an adjunct to MAR	99
III.6.a. Surgery prior to MAR in women with peritoneal endometriosis	99
III.6.b. Surgery prior to MAR in women with ovarian endometriosis	100
III.6.c. Surgery prior to MAR in women with deep endometriosis	101
III.7. Non-medical treatment strategies	103
III.8. Fertility Preservation	105
III.9 Impact of endometriosis on pregnancy and pregnancy outcome	107
III.9.a. Effect of pregnancy on endometriotic lesions	107
III.9.b. Possible complications during pregnancy from a pre-existing endometriosis lesion	108
III.9.c. Impact of endometriosis on early pregnancy (1 <sup>st</sup> trimester)	108
III.9.d. Impact of endometriosis on 2nd and 3rd trimester pregnancy and neonatal outcome	110
<b>IV. Endometriosis recurrence</b>	<b>114</b>
IV.1 Prevention of recurrence of endometriosis	114
IV.1.a. Surgical technique for prevention of recurrence	114
IV.1.b. Medical therapies for prevention of recurrence	115
IV.1.c. ART and endometriosis recurrence	118
IV.2 Treatment of recurrent endometriosis	120
IV.2.a. Medical treatment for recurrent endometriosis	120
IV.2.b. Surgical treatment of recurrent endometriosis	121
<b>V. Endometriosis and adolescence</b>	<b>123</b>
V.1. Diagnosis	123
V.1.a. Diagnostic process	123
V.1.b. Risk factors for adolescent endometriosis	123
V.1.c. Clinical symptoms	124
V.1.d. Clinical examination	125
V.1.e. Imaging	126
V.1.f. Laboratory parameters	127
V.1.f. Diagnostic laparoscopy	127
V.1.g. Histology	128
V.2. Treatment	130
V.2.a. Medical treatment	130
V.2.b. Surgical treatment	130
V.2.c. Combined medical and surgical treatment	133
V.3. Fertility preservation	135
<b>VI. Endometriosis and menopause</b>	<b>137</b>
VI.1. Endometriosis in postmenopausal women	137
VI.2. Treatment of endometriosis in postmenopausal women	139
VI.2.a. Surgical treatment	139
VI.2.b. Medical treatment	141
VI.3. Menopausal symptoms in women with a history of endometriosis	143
VI.3.a. MHT for menopausal symptoms in women with a history of endometriosis	143
VI.3.b. MHT and recurrence of endometriosis in women after natural menopause	143
VI.3.c. MHT and recurrence of endometriosis in women after surgical menopause	143
VI.3.d. MHT and risk of malignancy	144
VI.3.e. Regimen of MHT in women with a history of endometriosis	144
VI.4. Menopause-related major health concerns in women with endometriosis	147



<b>VII. Extrapelvic Endometriosis</b>	<b>148</b>
VII.1. Diagnosis	148
VII.1.a. Abdominal wall, umbilical, perineal and inguinal endometriosis	148
VII.1.b. Thoracic endometriosis	148
VII.2. Treatment	151
VII.2.a. Extrapelvic endometriosis of the abdominal wall, the umbilicus, and the inguinal region	151
VII.2.b. Thoracic and diaphragmatic endometriosis	151
<b>VIII. Asymptomatic endometriosis</b>	<b>154</b>
VIII.1. Treatment	154
VIII.2. Monitoring	154
<b>IX. Primary prevention of endometriosis</b>	<b>157</b>
IX.1. Risk factors and prevention	157
IX.2. Genetic predisposition	158
<b>X. Endometriosis and cancer</b>	<b>160</b>
X.1. Link between endometriosis and cancer	160
X.1.a. Somatic mutations	163
X.1.b. Impact of hormone treatments	163
X.2. Monitoring for detection of malignancy	167
X.3. Surgery and malignancy	169
<b>Annex 1: Guideline group</b>	<b>171</b>
Declarations of interest	176
<b>Annex 2: Abbreviations</b>	<b>178</b>
<b>Annex 3: Key Questions</b>	<b>180</b>
<b>Annex 4: Methodology</b>	<b>183</b>
Guideline development	183
Implications of recommendations	184
Strategy for review of the Guideline draft	185
Guideline implementation strategy	185
Schedule for updating the guideline	185
<b>Annex 5 : Flowcharts</b>	<b>186</b>
<b>Annex 6: Stakeholder review</b>	<b>191</b>
Annex 7 : Literature report	separate document
Annex 8 : Evidence tables and Evidence-to-Recommendations	separate document



→ PROGRAMME Thursday, 16<sup>th</sup> June 2022

6<sup>th</sup> EUROPEAN ENDOMETRIOSIS CONGRESS

June 16<sup>th</sup> & 17<sup>th</sup> 2022

BORDEAUX FRANCE Cité Mondiale www.eec2022.com

→ Thursday, June 16<sup>th</sup>

**Amphitheater BRISBANE**

08:00-08:30 **OPENING CEREMONY**

- 08:10 • The Congress President *Norine Rummen (France)*
- 08:20 • The President of EEL *Harald Brenner (Germany)*
- 08:30 • The Congress Secretary *Isabelle Charvatiat Lathrop (France)*

08:30-12:30 **LIVE SURGERIES**

*Moderators: Heidi von Elm (Germany), David Redwine (USA), Françoise Maillot (France), Michel Caron (France), Maurizio Giamberini (Italy), Prashant Mangalakar (India)*

- 09:00 • Surgery of sacral roots *Norine Rummen (France)*
- 09:30 • Rectal shaving using ICG *Norine Rummen (France)*

10:30-11:30 **Coffee Break**

- 11:00 • NDI-CECology for colorectal endometriosis *Andri Balazs (Hungary)*
- 11:30 • Ovarian endometriomas *Sergio Luis Mendez (France)*

12:15-12:30 **Live surgeries at the 5<sup>th</sup> EEC:**

- Patients postoperative outcomes and 2 year follow up *Jan Strahmowski (Czech Rep), Rachel Oswald (Czech Rep)*

12:30-13:30 **INDUSTRY SYMPOSIUM** (see detail p.23)

→ Thursday, June 16<sup>th</sup>

**Room BRASILIA**

08:30-10:30 **Fundamental science**

*Moderator: Silvia Mehnert (Germany)*

- Endometriosis-Associated Macrophages: origin, phenotype, and function *Enn Greenen (UK)*
- Problems and limitations of basic research in the field of endometriosis *Martin Götz (Germany)*
- Neuroimmunomodulation - clinical impact on endometriosis *Sylvia Mehnert (Germany)*
- Oral Communications *TBC*

10:30-11:00 **Coffee Break**

11:00-12:30 **Imaging Tips & Tricks**

*Moderators: Corinne Chénouet (Italy), Anneliina Mennonen-Peltonen (Finland)*

- RENDAN classification in ultrasound *Sarmin Madadi (Australia)*
- New classification of endometriosis using MRI *Isabelle Thomassin-Naggara (France)*
- What to look for: systematical analysis in 10 minutes *Frédéric Caronnet (France)*
- What looks like endometriosis and really is *Isabelle Thomassin-Naggara (France)*
- What looks like endometriosis and is not *TBC*

12:30-13:30 **INDUSTRY SYMPOSIUM** (see detail p.23)

→ PROGRAMME Thursday, 16<sup>th</sup> June 2022

6<sup>th</sup> EUROPEAN ENDOMETRIOSIS CONGRESS

June 16<sup>th</sup> & 17<sup>th</sup> 2022

BORDEAUX FRANCE Cité Mondiale www.eec2022.com

13:30-13:45 **Lunch Break**

**Amphitheater BRISBANE**

13:45-14:30 **DEBATE**

*Moderators: Heidi von Elm (Germany), Michel Caron (France)*

- Where does Endometriosis come from? Hypothesis & clinical consequences *David Redwine (USA), Dan C. Martin (USA)*
- Introduction to the debate: Is Endometriosis a progressive disease? *Michel Caron (France)*

14:30-14:45 **How to manage ovarian endometriomas?**

*Moderators: Françoise Maillot (France)*

- What imaging techniques? *Corinne Chénouet (Italy)*
- What medical treatment? *Isabelle Angelier (Italy)*
- What surgery? Debate: to excise, to ablate or sclerotherapy of ovarian endometriomas, and consequences on fertility:
  - excision: *Michel Caron (France)*
  - ablation: *Angela Dornes (Belgium)*
  - sclerotherapy: *Norine Rummen (France)*
- What about pregnancy and/or fertility preservation? *Miguel Ángel Soriano (Italy)*
- What short/long term outcomes? *Francesca Carmona (Spain)*

14:45-14:55 **Coffee Break**

13:30-13:45 **Lunch Break**

**Room BRASILIA**

13:45-15:15 **New insights about the medical treatments & guidelines in Europe**

*Moderators: Andrew Haines (UK), Lucy Whitaker (UK)*

- The evidence-base for medical management for endometriosis-associated pain *Anneliina Mennonen-Peltonen (Finland)*
- GnRH antagonists for endometriosis *Laura Ruggio (Italy)*
- Medical management to treat recurrence *Lucky Simoes (UK)*
- Progressive versus surgery for deep endometriosis *Federico Vercellotti (Italy)*
- Should progesterone be offered to all women with endometriosis-associated pain? *Isabelle Charvatiat Lathrop (France)*
- Novel therapeutic targets for endometriosis *Philippa Saunders (UK)*

15:15-16:00 **INDUSTRY SYMPOSIUM** (see detail p.23)

16:00-16:30 **Coffee Break**

→ PROGRAMME

Thursday,  
16<sup>th</sup> June  
2022

**6<sup>th</sup> EUROPEAN  
ENDOMETRIOSIS  
CONGRESS**

June  
16<sup>th</sup> & 17<sup>th</sup>  
2022

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Amphitheater BRISBANE

**14.30-15.00** **Honorary President lecture:**  
Recognition of endometriosis lesions  
*Den C. Martin (USA)*

**17.00-18.30** **SGP Session** (French Society of Pelvic Gynecological Surgery)

- My Journey with endometriosis  
*Christelle Zachariopoulos (France)*
- Is fibrosis a potential therapeutic target for endometriosis?  
*Sabrina Mousset (France)*
- Endofert: Lessons and Results  
*Fanny Carballat (France)*
- Endometriosis Organization of the research in France  
*Nicolas Bourde (France)*
- Epidemiology of Endometriosis in France Results of a national study  
*Sofiane Mial (France)*
- Ovarian vascularization assessment using fluorescence  
*Anna Sophie Goussard, Alexandre Jolyon (France)*

**18.30-19.30** • EEL General Assembly

**19.30-23.00 CONGRESS DINNER**

Room BRASILIA

**14.30-18.30** **Expert Center in endometriosis: what does it mean?**  
*Moderators: Louise Hummelshøj (Denmark), Marcello Casanovi (Italy)*

- In Denmark  
*Lone Hummelshøj (Denmark), Axel Forman (Denmark)*
- In France  
*Isabelle Chauvaud-Lynch (France), Yasmine Corbau (France)*
- In Germany and Austria  
*Norin Stein (Germany)*
- In Italy  
*Sofiane Angioni (Italy)*
- In Romania  
*Ioana Simedrea (Romania)*
- French regional endometriosis networks  
*Heide Roman (France)*
- Are policies indispensable to move forward?  
*Benoit Drouot (France)*
- EEL Certification EuroEndoCert  
*Berlin Renner (Germany)*
- Summary: Global consensus on the agreement on criteria for Expert Centers and individual certification  
*Lone Hummelshøj (Denmark)*

**19.30-23.00 CONGRESS DINNER**

→ PROGRAMME

Friday,  
17<sup>th</sup> June  
2022

**6<sup>th</sup> EUROPEAN  
ENDOMETRIOSIS  
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June  
16<sup>th</sup> & 17<sup>th</sup>  
2022

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→ Friday, June 17<sup>th</sup>

**08.00-09.20** **BEST VIDEOS**  
*Moderators: Euphémie Marlot (France), Sofiane Angioni (Italy)*

**09.20-10.00** **INDUSTRY SYMPOSIUM** (see detail p.23)

**10.00-11.00 Coffee Break**

**11.00-12.30** **How to manage a deep endometriosis of the digestive tract?**  
*Moderators: Carol Teoharova (USA), Heide Roman (France)*

- Which imaging techniques?  
*Isabelle Thomassin-Nagy (France)*
- Which surgical techniques?  
- shaving: *Clément Corinne (France)*  
- segmental resection: *Marcello Casanovi (Italy)*  
- disc excision: *Lucie Dang (France)*
- How to manage complications (natural/postoperative)?  
*Samuel Hubert (Australia)*
- If the patient wishes to get pregnant? What place for preoperative fertility preservation?  
*Sandra Courcier (France)*
- What about long term outcome?  
*Attila Bekes (Hungary)*

**13.30-13.00** **INDUSTRY SYMPOSIUM** (see detail p.23)

→ Friday, June 17<sup>th</sup>

**08.30-09.30** **ORAL COMMUNICATIONS**  
*Moderators: Louise Ulrich (Germany), Guendal Puelletre (France)*

**09.30-10.30** **Pain management**  
*Moderators: Denis Brusa (France), Jean-Luc Brun (France)*

- The world of pain of endometriosis : a plea for a global care  
*Arnoud Tausonier (France)*
- Endometriosis and central sensitization  
*Stephane Pothou (France)*
- From bench to bedside: a potential novel non-hormonal treatment for endometriosis-associated pain  
*Andrew Horne (UK)*
- From Gerring to Medical Approaches: Digital Therapeutics (DTx) in the Treatment of Endometriosis Pain  
*Serge Marchand (Canada)*

**10.30-11.00 Coffee Break**

**11.00-12.30** **Adenomyosis**  
*Moderator: Jacques Denon (Belgium), Lubov Mihaljevic (Czech Republic)*

- Imaging  
*Thibault Carteret (France)*
- HIFU and RFA  
*Claude Huet (France)*
- Hysteroscopic management: when and for whom?  
*Nicolas Fernandez (France)*

→ PROGRAMME

Friday,  
17<sup>th</sup> June  
2022

6<sup>th</sup> EUROPEAN  
ENDOMETRIOSIS  
CONGRESS

June  
16<sup>th</sup> & 17<sup>th</sup>  
2022

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Amphitheater BRISBANE

12:25-12:30 Lunch Break

13:30-14:00 Future opportunities in surgical management of complex deep endometriosis

Moderators: Axel Langer (Germany), Prasad Mangeshkar (India)

- Robotic surgery: does it provide any advantage?  
*Sally Stewart (UK)*
- Augmented reality in deep endometriosis: is it feasible?  
*Nicolas Bourde (France)*
- RENZIAN classification of endometriosis: a unique language  
*Jiang Weidong (Australia)*
- Finding Endometriosis using Machine Learning: A bird's-eye view of the FEMALE project  
*Lene Bak Kirk (Denmark)*

14:30-14:40 ART and surgery: let's talk about infertility management

Moderators: Elina Aho (France)

- Infertility explained to surgeons: what they should know about ART?  
*Isabelle Charrier Lehericy (France), Sandrine Courbiere (France)*
- Surgery explained to ART specialists: what they should know about surgery?  
*Norica Roman (France), Marcus Sebastian (France)*

Room BRASILIA

- Infertility risk  
*Louis Marzolin (France)*
- Obstetrical risk  
*Rache Versalini (Italy)*
- Is surgery worthwhile? What place for Ovaria procedure?  
*Harald Krentz (Germany)*

12:30-12:30 Lunch Break

13:30-14:15 Endometriosis and infertility: questions about AMH

Moderators: Isabelle Charrier Lehericy (France)

- How to trust AMH?  
*Oskar Dewailly (France)*
- Should we prescribe AMH to every patient presenting with endometriosis?  
*Erkin Oral (Turkey)*
- Gold standard for decision of fertility preservation?  
*Sandrine Courbiere (France)*

14:15-14:40 Requirements for education in endometriosis - the residents' view

Moderators: Andrea Nappo (Germany), Gulnur Topcu (Turkey), Ann-E. Esslinger (Germany)

→ PROGRAMME

Friday,  
17<sup>th</sup> June  
2022

6<sup>th</sup> EUROPEAN  
ENDOMETRIOSIS  
CONGRESS

June  
16<sup>th</sup> & 17<sup>th</sup>  
2022

BORDEAUX - FRANCE  
Cité Mondiale  
[www.endometriosis.org](http://www.endometriosis.org)

Amphitheater BRISBANE

14:30-14:30 Coffee Break

16:30-18:00 How to manage a deep endometriosis of the sacral plexus?

Moderators: Lucie C. MATHIEU (UK), Aditi Arora (Singapore)

- Anatomy of the parametrium. Symptoms of endometriosis involving sacral plexus  
*Michael Seyer Hansen (Denmark)*
- What preoperative assessment?  
*Shahen Khachat (UK)*
- Nerve sparing vs. radicality: is radicality always necessary?  
*Marcello Di Giacomo (Italy)*
- Excision of deep nodules of parametria in 10 steps  
*Norica Roman (France)*
- Neuropraxia, what does it mean?  
*Prashant Mangeshkar (India)*
- What short/long term outcomes?  
*Steph Norman (Denmark)*

18:00 Closing Ceremony

Room BRASILIA

14:45-16:00 Endometriosis and cancer

Moderators: Erwin Dahan (France), Aditi Arora (Singapore)

- Endometriosis and cancer -  
*Murina Kusdoff (France)*
- How to manage large ovarian endometriosis in premenopausal patients? Oophorectomy or not?  
*Aditi Arora (Singapore)*
- Risk of cancer - reasons for radical surgery in symptom free patients?  
*Olivia Bratta (Romania)*
- Hormon replacement therapy and risk of malignant transformation of endometriosis  
*Erwin Samliger (UK)*

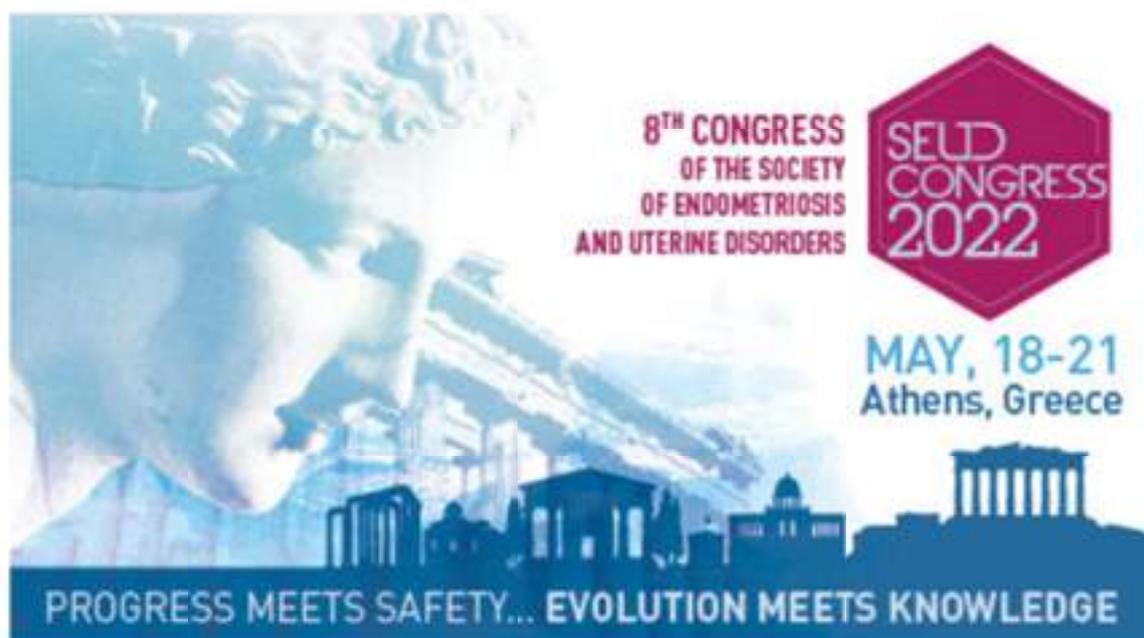
14:45-14:45 Coffee Break

16:30-18:00 Early-Career WES session:

Moderators: Lucy Whicker (UK), Erwin Dahan (UK)

- Career path testimonials  
*Philippa Saunders (UK), Michel Carré (France), Harald Krentz (Germany)*
- Winners programme  
*Sebastian Schäfer (Germany)*
- WES mentoring programme  
*Murina Kusdoff (France)*

**SEUD 2022**



The 8<sup>th</sup> Congress of The Society of Endometriosis and Uterine Disorders will be held in May in Athens.

## D INTERVIEW WITH AN 'ENDO SPECIALIST'



**Mario Malzoni**  
Interviewer: Ezgi Darici

**Turkish Endometriosis and Adenomyosis Society (EAD):** Doctor Malzoni, thank you very much for accepting our invitation to this interview. You have been dealing with endometriosis for many years. Can you tell us how your journey with endometriosis started?

**Mario Malzoni (MM):** My experience with endometriosis began many years ago in 1995. After my education at the University of Naples, I went to Columbia University in New York, where I worked with Professor Harry Reich. Harry Reich is one of the most important pioneers in the research and surgical treatment of endometriosis and especially in the laparoscopic and surgical treatment of deep endometriosis. I started working with my mentor Harry Reich in the US in 1996 and spent a year with him at both Columbia University and a private centre, and then I came back to my home centre in Italy. My clinic is a private clinic, but it has an agreement with the Italian healthcare system. So, we operate like a public hospital in a very large geographical area in the southern part of Italy, so we have many patients with deep endometriosis. In 1997, we started operating on difficult endometriosis cases, a general surgeon was working in my clinic at that time.

Then I started to assemble a team of young doctors specializing only in laparoscopic surgery, which included gynaecology, oncology and deep endometriosis surgeons, but no obstetricians. Then we started operating on cases of bowel and bladder endometriosis. We're currently doing at least 800-900 surgeries a year at The Malzoni Centre. Our current philosophy is that the gynaecology team should do all the work during the operation themselves. So, we do everything from bowel surgery to bladder and ureter surgery which is very important. Because in this way, you and your entire team both increase your experience and improve in the management of complications.

**EAD: What do you think is the most challenging aspect of the disease? Diagnosis or treatment?**

**MM:** I think every aspect of deep endometriosis is challenging. The challenge starts from the diagnosis. Sometimes the diagnosis of the disease is very difficult, and we need a very good diagnostic capability. There is a very good ultrasonographer in our group. Dr. DiGiovanni is a person with very high diagnostic accuracy and specializes in the diagnosis of deep endometriosis. We have published all our data in our clinic. At the level of the posterior compartment, lateral compartment, bowel, bladder, anterior compartment, ureter, and parametria, we have a diagnostic accuracy of greater than 90 percent, sometimes 100 percent. This is really important. And if you evaluate patients very well at the beginning, you can plan the right strategy which sometimes means medical treatment, pregnancy, or sometimes surgery. When the surgery decision is made, if the correct diagnosis is made, you can talk to the patients about the surgical strategy, the complications that may occur during the surgery, and therefore you will not encounter any surprises during the surgery. In this case, everything is planned before entering the operating room. So if you need bowel preparation for bowel surgery, you have time, or you need cooperation with other doctors, general surgeon or urologist, you can be better organized before surgery, reducing the risk of surprises and of course reducing the risk of complications. The biggest challenge of endometriosis surgery is deep endometriosis, of course, in my opinion, bowel surgery, bladder and ureter surgery but especially parametrial surgery. Surgery becomes very difficult at the level of the parametrium, especially in the presence of bilateral parametrial involvement, especially in the dorsal and lateral parametrial involvement. This is because your risk of denervation increases. You should explain this situation to the patient, share all the risks with the patient, and plan the right surgery, especially for young patients, because your risk of neurological complications will be high after such a surgery. While removing the lesions, you must perform the surgery without damaging the nerves. For this reason, I think parametrial endometriosis is the most difficult part of the surgery.

**EAD: You see a lot of endometriosis patients. Some of your patients are adolescent and perimenopausal patients who are at the beginning and end of their fertility periods. Do you have different strategies for the management of these patients?**

**MM:** Yes, of course. For me, the first try is definitely medical treatment. Therefore, if we have an ultrasound evaluation suggestive of deep endometriosis in adolescent patients, of course, we sometimes consider the option of surgery, but you know that the prevalence of deep endometriosis in adolescent patients is very low. That is, there is usually superficial-peritoneal endometriosis. In my opinion, the necessity of surgery in adolescent patients is rare. However, if surgery is necessary, you need to implement medical treatment after the surgery. Therefore, I usually prefer to start with medical treatment. Of course, you need long-term medical treatment. For this reason, initially you may be able to start with a progesterone-containing drug, possibly dienogest, followed by continuous oestrogen-progesterone drugs. Strategy in perimenopause depends on diagnosis. If there is a case of deep endometriosis with infiltration of organs and you suspect organ damage, such as the dilatation of bladder, ureter, or intestinal obstruction, then of course, surgery is required. But if there is no organ involvement and the patient is of perimenopausal age, medical treatment is likely to work well, and you can continue with medical treatment until the age of menopause. If medical treatment doesn't work and if adenomyosis is present with deep endometriosis, you can of course decide to remove the uterus and ovaries and take a radical procedure to completely stop oestrogen production.

**EAD: You are one of the pioneers in the field of endometriosis surgery. Can you give us some tips on how we can prevent complications during surgery? Of course, you mentioned pre-operative mapping before, but what if something happens during surgery?**

**MM:** Yes, if we do a good mapping before the surgery, the risk of surprise is very low, but of course you always have the risk for complications. So, I think many of the tips and tricks you need to reduce the risk of complications are based on a good surgical strategy. So, to get to the disease, search the retroperitoneal anatomy starting from the normal tissue, see the ureters, see the nerves, know the nerve anatomy so you know exactly where the nerves are and then you can go inside the disease to remove the disease. The use of electricity, bipolar, monopolar is very important. The use of ultrasound devices is very important. Circular stapler use and technology knowledge are very important in intestinal surgery. We have started to use diode laser in the treatment of superficial endometriosis, and especially in the treatment of endometrioma, to reduce the damage to the ovarian tissue. We use the laser to avoid excision and evaporate the capsule, especially in cases of bilateral endometrioma. We know that the risk of recurrence is probably higher compared to excision, but I think it is a better method of reducing the risk of ovarian damage, especially if a young woman has bilateral endometrioma.

**EAD: You told us about your endometriosis journey and said that you have worked with many young doctors and surgeons. Do you have any advice for young colleagues who want to specialize in endometriosis?**

**MM:** Sure. The recommendation is that if you want to do this type of surgery, you have to completely lose your obstetric side. You just have to focus on gynaecology. And, of course, you need to be well-trained in a specialized centre for endometriosis (excellence centre) and work collaboratively with radiologists or sonographers to compare the diagnosis and be in constant communication every day. To compare the diagnosis with intraoperative evaluation, you need to spend all day in the operating room with a trained sonographer or radiologist.

**EAD: It is very nice that you not only do your daily practice but also organize Malzoni Meetings. How did you start organizing these meetings?**

**MM:** Since the number of doctors specializing in the treatment of deep endometriosis is very few, we set out with the idea of sharing our experiences with our colleagues in order to share scientific knowledge and increase the number of experts in this field. Thus, we started organizing meetings in our own centre in 1999. Every year we organize a big meeting, national, international, many courses throughout the year. We organize different courses for oncology, endometriosis, up-to-date technological approach courses on pelvic, and we hold our international endometriosis meeting every three years. We did the last one online due to COVID and now we will have a very big meeting in Rome in 2023. In May, we will organize it together with AAGL, European Society of Gynaecology and Italian Society of Gynaecological Endoscopy. During the four-day congress not only endometriosis, but also oncology, benign diseases and hysteroscopy will be discussed. We have also developed a new platform, 'Malzoni TV', based on our COVID experiences, and it continues very well. At least 1000 people access the platform every Friday. We decided to organize one Friday every month, this event is called 'Friday I am Online'. We organize the event with different topics every month.

**EAD: And my last question is, do you offer fellowship opportunities at your centre?**

**MM:** Yes, of course. We have many residents and fellows from different parts of the world here. We have at least two people from other countries and two assistants from the Italian University. In other words, four people are with us every day, and the duration of their participation can be three months, six months, one year, sometimes up to two years.

**EAD: Thank you very much for your time and for your valuable contribution.**

## E ARTICLES ON ENDOMETRIOSIS FROM OUR COUNTRY FROM THE LAST THREE MONTHS

### 1. Effects of ranibizumab and zoledronic acid on endometriosis in a rat model.

Ureyen Ozdemir E, Adali E, Islimye Taskin M, Yavasoglu A, Aktug H, Oltulu F, Inceboz U.

Arch Gynecol Obstet. 2022 Jan;305(1):267-274. doi: 10.1007/s00404-021-06104-9. Epub 2021 Jun 3. PMID: 34081204.

**Purpose:** To investigate the histological efficacy of ranibizumab and zoledronic acid in an experimentally induced endometriosis model as compared with danazol, buserelin acetate and dienogest.

**Methods:** Endometrial implants were introduced in 52 female Wistar albino rats, which were then randomly divided into six groups. The animals were, respectively, given dienogest, danazol, buserelin acetate, zoledronic acid, ranibizumab and 0.9% NaCl. After 4 weeks, the volumes and histopathological properties of the implants were evaluated and the implants were excised completely at the third laparotomy. A histopathological scoring system was used to evaluate the preservation of epithelia. Endometrial explants were evaluated immunohistochemically.

**Results:** Among the groups, the histological score was significantly lower in the zoledronic acid and ranibizumab groups compared with the controls ( $p < 0.001$ ). There were no significant differences regarding ellipsoidal volume levels between groups ( $p > 0.05$ ). However, there was a statistically significant difference regarding cell numbers according to the degree of Bcl-2, NF- $\kappa$ B, and CD31 staining ( $p < 0.001$ ). There was no statistically significant difference in Bcl-2, CD31, or NF- $\kappa$ B staining in the binary comparisons between the other groups ( $p > 0.05$ ). For Bcl-2 staining, the staining rate of the group treated with zoledronic acid was significantly lower compared with the dienogest and danazol groups ( $p < 0.05$ ). The staining rates of CD31 and NF- $\kappa$ B were significantly lower in the zoledronic acid and ranibizumab groups compared with the controls ( $p < 0.05$ ).

**Conclusion:** According to these results, zoledronic acid and ranibizumab may be putative candidates for the treatment of endometriosis.

### 2. T-Cadherin, E-Cadherin, PR- $\alpha$ , and ER- $\alpha$ Levels in Deep Infiltrating Endometriosis

Kalkan U, Biyik I, Simsek S. Int J Gynecol Pathol. 2022 Feb 11. doi: 10.1097/PGP.0000000000000860. Epub ahead of print. PMID: 35149616.

#### Abstract

The goal of this study was to compare the T-cadherin, E-cadherin, progesterone receptor (PR), and estrogen receptor (ER) staining levels of deep infiltrating endometriosis (DIE) tissue, ovarian endometriomas and normal endometrial tissues in the same individuals. The tissue sections of both DIE nodule(s) and endometrioma(s) of 15 cases were examined. As a control group, normal endometrial tissue sections of 23 cases were examined. T-cadherin, E-cadherin, ER- $\alpha$ , and PR- $\alpha$  staining levels of DIE, endometrioma tissues, and endometrial tissues were compared immunohistochemically. H-score was calculated to compare the expression of T-cadherin, E-cadherin, ER- $\alpha$ , and PR- $\alpha$  in immunohistochemical staining based on the percentage of cells stained at each intensity level. T-cadherin, E-cadherin, ER, and PR H-score were lowest in DIE tissue and highest in endometrial tissue ( $P < 0.0001$ ,  $< 0.0001$ ,  $< 0.0001$ , and  $< 0.0001$ , respectively). In correlation analysis, a positive correlation was found between T-cadherin, E-cadherin, PR, and ER H-score ( $P < 0.0001$  for each). T-cadherin, E-cadherin, ER, and PR H-score were lowest in DIE tissue and highest in endometrium tissue. We think that examination of DIE tissue and endometrioma tissue from the same individual excludes the possibility of an effect due to different genetic and environmental factors from different individuals. With the help of this exclusion we showed that DIE and endometrioma have different biological properties.

### 3. Analysis of Changes in Serum Levels and Gene Expression Profiles of Novel Adipocytokines (Omentin, Vaspin, Irisin and Visfatin) and Their Correlation with Serum C-reactive Protein Levels in Women Diagnosed with Endometriosis.

Kaya Sezginer E, Kirlangıç ÖF, Eşkin Tanrıverdi MD, Topçu HO, Gür S. Turk J Pharm Sci. 2022 Feb 28;19(1):48-53. doi: 10.4274/tjps.galenos.2021.52284. PMID: 35227049; PMCID: PMC8892558.

**Objectives:** This study aimed to investigate the role of new adipocytokines (omentin, vaspin, irisin and visfatin) in the development of endometriosis and the relationship of these adipocytokines with the inflammatory marker, C-reactive protein (CRP) levels in serum.

**Materials and methods:** In this study, endometriosis (n=16) and control groups (n=14) were determined *via* ultrasound. Serum omentin, vaspin and irisin levels were measured by ELISA method. CRP levels in serum and the gene expression of visfatin and vaspin in whole blood samples were determined by clinical analyzer and the real-time polymerase chain reaction, respectively.

**Results:** Serum irisin and CRP levels in the endometriosis group were significantly higher than in the control group. Irisin protein levels demonstrated a positive correlation with body mass index and CRP in women diagnosed with endometriosis. No statistically significant difference was found in serum omentin and vaspin levels between groups. The visfatin and vaspin gene expression in whole blood samples from the endometriosis group was found to be significantly lower than the control group.

**Conclusion:** Increased levels of serum irisin and decreased visfatin and vaspin gene expressions in blood may be considered as a potential biomarker in endometriosis. The identification of new adipocytokines, which demonstrate an alteration in the presence of endometriosis and the relationship between these adipocytokines and inflammation will facilitate the detection of mechanisms involved in endometriosis and will lead to the development of targeted therapy.

#### 4. The effects of adalimumab on the rat autotransplantation endometriosis model: A placebo-controlled randomized study.

Kaplan S, Kırıcı P, Türk A. *Adv Clin Exp Med*. 2022 Jan 18. doi: 10.17219/acem/144369. Epub ahead of print. PMID: 35040600.

**Background:** Endometriosis is a chronic inflammatory pathology that can cause persistent pelvic pain and infertility by affecting women of reproductive age. It is defined as the placement of endometrial tissue outside the uterine cavity. Hormonal, genetic and immunological factors have an effect on the development of endometriotic implants. Adalimumab is a monoclonal antibody specific for tumor necrosis factor alpha (TNF- $\alpha$ ), used in the treatment of autoimmune diseases.

**Objectives:** To investigate the effectiveness of adalimumab on histopathological and biochemical values in rats with experimental endometriosis.

**Material and methods:** This study is a comparative, prospective, experimental rat study. Wistar albino female rats were divided into 4 groups. Group 1 was separated as the control group. Endometriotic implants were simultaneously induced in group 2 and group 3. After 4 weeks, developing endometriotic foci were measured. Adalimumab (5 mg/kg) was simultaneously intraperitoneally (ip.) administered to group 3 and group 4 for 4 weeks. At the end of the study, histopathological scoring and fibrillin-1 scoring were performed. Total antioxidant status (TAS), total oxidant status (TOS) and malondialdehyde (MDA) values were measured. Findings in all groups were compared.

**Results:** When group 1 and group 2 were compared, the histopathological score, as well as MDA and TOS levels increased, while TAS levels decreased in group 2 ( $p < 0.001$ ). After adalimumab treatment, the average endometriotic implant size in group 3 ( $0.32 \pm 0.002$  mm) decreased compared to group 2 ( $0.77 \pm 0.04$  mm) ( $p = 0.032$ ). While fibrillin-1 score increased in group 2 and group 3 compared to group 1, it decreased in group 3 compared to group 2 ( $p < 0.001$ ). Histopathological score decreased, TAS levels increased and MDA levels decreased in group 3 compared to group 2 ( $p < 0.001$ ).

**Conclusions:** Adalimumab may play a role in the regression of endometrial implants by showing antioxidant and anti-inflammatory effects on histopathological damage and fibrosis.

#### 5. Predictive value of preoperative MRI using the #ENZIAN classification score in patients with deep infiltrating endometriosis.

Fendal Tunca A, Iliman DE, Akdogan Gemici A, Kaya C. *Arch Gynecol Obstet*. 2022 Mar 3. doi: 10.1007/s00404-022-06451-1. Epub ahead of print. PMID: 35239004.

**Purpose:** The aim of this study is to investigate the correlation between the magnetic resonance imaging (MRI) and intraoperative findings of deep infiltrating endometriosis using the #ENZIAN score.

**Methods:** This retrospective study included 64 patients who underwent surgery for deep infiltrating endometriosis between January 2017 and August 2020. Preoperative abdominopelvic MRI assessment was evaluated and scored using the #ENZIAN classification. Operative scores were considered the gold standard, and the sensitivity, specificity, and positive and negative predictive values (PPV and NPV) of MRI for each category were calculated.

**Results:** MRI has higher sensitivity and specificity in showing the lesions of the compartments O (ovarian lesions), A (rectovaginal septum and posterior vaginal fornix), and B (uterosacral ligaments and parametrium) (100-100%, 100-100%, and 97-100%, respectively,  $p < 0.001$ ) compared to the other compartments. The lowest sensitivity, specificity, accuracy, and PPV of the MRI was found in compartment P (14%, 76%, 70%, and 7%, respectively).

**Conclusion:** We demonstrated that the #ENZIAN classification in MRI reports has significant sensitivity and specificity in compartments A, B (uterosacral ligaments and parametrium), and O. Furthermore, the determination of peritoneal lesions via MRI is inadequate.

### 6. The rate of oocytes with granular cytoplasm is higher in women with endometrioma in ICSI cycles.

Bilgic BE, Kurek Eken M, Ayla Ş, Kose A, Kutlu T, İlhan G. J Obstet Gynaecol. 2022 Apr;42(3):467-471. doi: 10.1080/01443615.2021.1916803. Epub 2021 Jun 24. PMID: 34165007.

#### Abstract

The purpose of this study was to investigate the impact of endometrioma on oocyte morphology and fertility outcome in intracytoplasmic sperm injection (ICSI) cycles. The study material was obtained from 114 ICSI cycles of infertile women aged between 20 and 38 years with ovarian endometriomas and unexplained infertility. In total, 644 mature oocytes were included in the analysis. The rates of specific oocyte morphological abnormalities were similar between the two groups however the central granulation rate was significantly higher in the group with endometrioma ( $p < .05$ ). Fertilisation rate were not significantly different between the groups ( $p \geq .05$ ) however the numbers of metaphase 2 (MII) oocytes and embryos were lower in the endometrioma group ( $p \leq .05$ ). Endometrioma was associated with a higher rate of oocytes with granular cytoplasm, despite the fertilisation rate the numbers of the MII oocytes and embryo were affected. IMPACT STATEMENT

**What is already known on this subject?** The association between endometrioma and infertility is a well-known condition, but the possible mechanisms of the effects of endometrioma on women's fertility is still debated and controversial. There is limited data on the effect of endometrioma on oocyte morphology. Low oocyte quality and lower fertilisation rates might be the main cause of adverse pregnancy outcomes during *in vitro* fertilisation/intracytoplasmic sperm injection cycles.

**What do the results of this study add?** Endometrioma was associated with a higher rate of oocytes with granular cytoplasm, and lower metaphase 2 oocytes and embryos.

**What are the implications of these findings for clinical practice and/or further research?** Future studies using further oocyte quality assessment methods and prospective observational studies including live-birth rate should be designed to better understand how endometrioma affects fertility outcomes.

### 7. Bisphenol A levels in bowel endometrioma diagnosed serums: A case control study.

Ardic F, Celik H, Yesilyurt H, & Ozcelik Otcu S. EXPERIMENTAL BIOMEDICAL RESEARCH, 5(1), 38-47. <https://doi.org/https://doi.org/10.30714/j-ebr.2022173849>

**Aim:** To investigate the bisphenol A (BPA) levels, which may be a risk factor in the etiology of endometrioma, in patients diagnosed laparoscopically with endometrioma with and without bowel involvement.

**Method:** In the prospective cross-sectional case control study, 47 cases were included in the study, which were admitted to the gynecology and infertility services with and without bowel involvement endometrioma who were operated and diagnosed histopathologically. 43 patients were included in the control group. For serum BPA value, blood samples taken immediately before the operation were studied in laboratory. Patients and controls were compared with controls in terms of serum BPA values.

**Results:** The mean age of the patients was  $35 \pm 2$  in the endometriosis group and  $36 \pm 2$  in the control group which was and not statistically significant. There was no statistical difference between the patient and control groups in terms of menstruation periods. Serum BPA levels were significantly higher in the bowel involvement group compared to the non-bowel involvement group, as the distribution width was higher due to excessive values, and only 5 patients with bowel involvement did not reach statistically significant levels. Serum BPA level was  $1084 \pm 1132$  ng/L in the endometriosis group and  $269 \pm 99$  ng/L in the control group which was statistically significant ( $p < 0,001$ ).

**Conclusions:** BPA levels were showing very wide range especially in the patient group. Serum BPA levels was statistically significantly higher in the endometrioma group compared to the control group. Therefore, in the etiology of endometriosis BPA may take a definite place.

### 8. A Rare Cause of Bowel Obstruction Mimicking Colon Cancer: Endometriosis.

Açar S, Çiftçi E, Çetiner H, Api M. J Clin Obstet Gynecol. 0;0(0):0-0.

#### Abstract

Intestinal involvement is observed in a significant portion of patients diagnosed with endometriosis. Involvement is often on the serosal surface of the sigmoid colon. Rarely, it takes place in the colonic mucosa and lymph nodes, causing lumen obstruction. Colonoscopic biopsies may be insufficient in the differential diagnosis of malignancy and diverticulitis. We report on a case of a 41-year-old premenopausal female patient endometriosis located in the ampulla recti causing complete intestinal obstruction was detected. Although having a significant rectal mucosal mass, the diagnosis could not be made with tissue samples taken by endoscopy. During the examination, the patient developed ileus. With the preliminary diagnosis of rectal cancer, surgical intervention was applied with oncological principles, and its treatment was performed with opening a diverting ileostomy. Diagnosis of rectosigmoid endometriosis is difficult. In women of childbearing age, rectosigmoid endometriosis should be kept in mind in lower gastrointestinal tract obstructions.

### **9 . Abdominal wall endometriosis: Case-series study and a systematic review.**

Katırcı Y. , Özdemir A. , Gülümser Ç. , Gun S. Journal of Experimental and Clinical Medicine. 2022; 39(1): 292-297.

#### **Abstract**

Abdominal wall endometriosis (AWE) is the presence of endometrial gland and stroma in the abdominal wall that should be kept in mind in differential diagnosis of pelvic pain due to especially increased C-section rates. Between January 2000 and July 2018, MEDLINE and EMBASE databases were systematically reviewed using the search criteria "abdominal wall endometriosis," "abdominal wall endometriomas,". Only the studies having over 20 patients were included. Case-series, case-control studies, and articles in languages other than English were excluded. Number of the patients, patients 'age, study design, previous surgical history, most common symptom, time interval to symptoms, treatment, recurrence rate, and tumour size were investigated. In Total, 18 studies and 994 women were included in the study. Case studies, studies with less than 20 cases, non-English articles were excluded from the study. In the included studies, the numbers of minimum and maximum woman were 20 and 227, respectively. AWE significantly impairs the quality of life in reproductive age patients and is commonly seen in women with previous history of laparotomy, especially those who underwent cesarean section. Therefore, it must be kept in mind in the differential diagnosis of women who have a history of pain and a history of previous surgery.

### **10. A DESMOID TUMOR IN PREGNANCY MIMICKING SUBCUTANEOUS ENDOMETRIOSIS; A CASE REPORT.**

Ertekin, Arif Aktug, et al.Ertekin, Arif Aktug, et al. Journal of Istanbul Faculty of Medicine, vol. 85, no. 1, Mar. 2022, pp. 136+. Gale OneFile: Health and Medicine, link.gale.com/apps/doc/A691769379/HRCA?u=anon~108b5c25&sid=googleScholar&xid=cb223046. Accessed 15 Mar. 2022.

#### **Abstract**

Desmoid tumors are benign non-metastasizing tumors. However, they may proliferate and infiltrate into adjacent tissues with high recurrence rates. Even though its etiology is unclear, a previous histories of trauma, surgery, pregnancy, use of medication containing estrogen, and having conditions such as familial adenomatous polyposis and Gardner syndrome can be regarded as contributing or risk factors.

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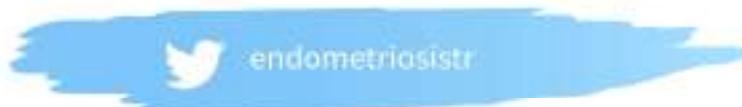
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