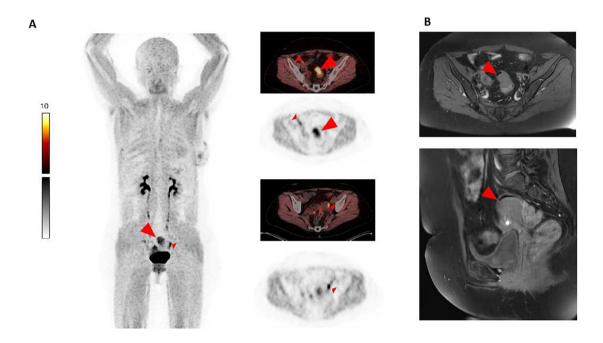
## **IMAGE OF THE MONTH**



## [<sup>68</sup>Ga]Ga-FAPI-04 PET/CT in a patient with endometriosis: a potential game changer?

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A 43-year-old woman with a known history of endometriosis presented with dysmenorrhea, dyschezia, and hematochezia. She had previously undergone a hysterectomy, during which infiltrative endometriosis in the bowel was

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identified. To guide further treatment decisions, a magnet resonance imaging (MRI) was performed, confirming bowel involvement and revealing a circumscribed lesion infiltrating the sigmoid colon. Due to persistent symptoms, the patient opted for surgery. For preoperative planning, a fibroblast activation protein inhibitor (FAPI) positron emission tomography/ computed tomography with [68Ga]Ga-FAPI-04 was conducted. PET images, acquired 10 min after administration 148 MBq of [68Ga]Ga-FAPI-04, showed intense tracer uptake (Fig. A; left column: maximum intensity projection; right column: axial slices of PET/CT fusion and PET; big arrow head: endometriosis in the sigmoid colon, SUVmax 9.8) at the same location as the MRI finding (Fig. B, axial and sagittal slices of contrast enhanced T1-weighted fat saturated sequence), thus confirming the MRI result. Additionally, the PET scan revealed intense uptake in the left



ovary and the ligamentum teres uteri bilaterally (Fig. A, small arrowheads; SUVmax: 11.3, 7.3 and 5.4, respectively) suggesting the presence of additional endometriotic lesions undetected by MRI. The patient underwent a second surgical procedure, which histopathologically confirmed a deep infiltrating endometriosis extragenitalis.

Endometriosis, characterized by the ectopic growth of endometrial-like tissue, is a common gynecological disease affecting approximately 10% of reproductive-age women [1]. It can cause significant symptoms, including dysmenorrhea, dyspareunia, infertility, and gastrointestinal disturbances [2]. In addition to its physical effects, endometriosis can also affect mental health [3]. Thus, endometriosis significantly impact the quality of life of patients [4]. Diagnosis is primarily symptom-based, which can be challenging due to their variety and the heterogenic phenotype of endometriosis overlapping with other diseases. A definitive diagnosis is mostly confirmed through diagnostic laparoscopy. While transvaginal ultrasound is the first-line imaging modality, MRI is often used as a second-line tool for assessing disease extent and aiding in surgical planning [5].

This case highlights the potential role of <sup>68</sup>Ga-FAPI PET/ CT in detecting endometriotic lesions. Not only did the scan confirm the known lesion, but also identified additional sites of involvement undetected by MRI. Many studies have demonstrated the biodistribution of the <sup>68</sup>Ga-labeled FAPI tracer, often showing intense uptake in the uterus [6-8]. There has been a previous case report showing only faint uptake in one endometriotic lesion [9]. To the best of our knowledge, this is the first confirmed case of endometriosis demonstrated on <sup>68</sup>Ga-FAPI PET/CT with clear and intensive uptake. Therefore, <sup>68</sup>Ga-FAPI PET/CT could represent a promising diagnostic modality for endometriosis, alongside many malignant diseases in which it has proven effective [10–13]. Our observation is supported by a recent preclinical study that found FAP expression in the immune microenvironment of endometriosis from various locations [14].

This case may provide a rationale for future studies systematically analyzing FAP expression in endometriosis across larger patient cohorts, investigating the value of <sup>68</sup>Ga-FAPI PET/CT. A prospective study was recently initiated and is currently underway investigating this issue (NCT06792318).

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**Data availability** The data that support the findings of this study are available from the corresponding author upon reasonable request.

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## References

- Viganò P, Parazzini F, Somigliana E, Vercellini P. Endometriosis: epidemiology and aetiological factors. Best Pract Res Clin Obstet Gynaecol. 2004;18:177–200.
- Zondervan KT, Becker CM, Missmer SA. Endometriosis. N Engl J Med. 2020;382:1244–56.
- Gambadauro P, Carli V, Hadlaczky G. Depressive symptoms among women with endometriosis: a systematic review and metaanalysis. Am J Obstet Gynecol. 2019;220:230–41.
- Gete DG, Doust J, Mortlock S, Montgomery G, Mishra GD. Impact of endometriosis on women's health-related quality of life: A national prospective cohort study. Maturitas. 2023;174:1–7.
- Quesada J, Härmä K, Reid S, Rao T, Lo G, Yang N, et al. Endometriosis: A multimodal imaging review. Eur J of Radiol. 2023;158: 110610.
- Giesel FL, Kratochwil C, Lindner T, Marschalek MM, Loktev A, Lehnert W, et al. <sup>68</sup>Ga-FAPI PET/CT: Biodistribution and Preliminary Dosimetry Estimate of 2 DOTA-Containing FAP-Targeting Agents in Patients with Various Cancers. J Nucl Med. 2019;60:386–92.
- Kessler L, Ferdinandus J, Hirmas N, Zarrad F, Nader M, Kersting D, et al. Pitfalls and Common Findings in <sup>68</sup>Ga-FAPI PET: A Pictorial Analysis. J Nucl Med. 2022;63:890–6.
- Zhang X, Song W, Qin C, Song Y, Liu F, Hu F, et al. Uterine Uptake of <sup>68</sup>Ga-FAPI-04 in Uterine Pathology and Physiology. Clin Nucl Med. 2022;47:7–13.
- Zou Y, Hu H, Tian R, Wang R. Focal <sup>68</sup>Ga-FAPI-04 Uptake in Endometriosis at the Rectouterine Pouch. Clin Nucl Med. 2025;
- Kratochwil C, Flechsig P, Lindner T, Abderrahim L, Altmann A, Mier W, et al. <sup>68</sup>Ga-FAPI PET/CT: Tracer Uptake in 28 Different Kinds of Cancer. J Nucl Med. 2019;60:801–5.
- Mori Y, Dendl K, Cardinale J, Kratochwil C, Giesel FL, Haberkorn U. FAPI PET: Fibroblast Activation Protein Inhibitor Use in Oncologic and Nononcologic Disease. Radiology. 2023;306: e220749.
- Dendl K, Koerber SA, Finck R, Mokoala KMG, Staudinger F, Schillings L, et al. <sup>68</sup>Ga-FAPI-PET/CT in patients with various gynecological malignancies. Eur J Nucl Med Mol Imaging. 2021;48:4089–100.
- Wegen S, van Heek L, Linde P, Claus K, Akuamoa-Boateng D, Baues C, et al. Head-to-Head Comparison of [<sup>68</sup> Ga]Ga-FAPI-46-PET/CT and [<sup>18</sup>F]F-FDG-PET/CT for Radiotherapy Planning in Head and Neck Cancer. Mol Imaging Biol. 2022;24:986–94.
- Kellers F, Lützen U, Verburg F, Lebenatus A, Tesch K, Yalcin F, et al. FAP-positive activated fibroblasts are detectable in the microenvironment of endometriosis and correlate with stroma composition and infiltrating CD8+ and CD68+ cells. Hum Reprod Open. 2025;hoaf003.

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