

Clinical Policy: Fertility Preservation

Reference Number: PA.CP.MP.130

Effective Date: 11/18

Date of Last Revision: 05/23

Coding Implications

Revision Log

Description

Fertility may be transiently or permanently affected by medical treatments such as gonadotoxic therapy, cytotoxic chemotherapy, or radiation therapy, as well as by other iatrogenic causes. Rates of permanent infertility and compromised fertility after medical treatment vary and depend on many factors, including the drug, size, and location of the radiation field if applicable, dose, dose-intensity, method of administration (oral versus intravenous), disease, age, treatment type and dosages, and pretreatment fertility.

Policy/Criteria

- I. It is the policy of Pennsylvania Health and Wellness® (PHW) that, when a covered benefit under the member's/enrollee's benefit plan contract, any of the following procedures are medically necessary for adults and adolescents with a female reproductive system prior to commencing medically necessary treatment that is likely to cause infertility (excluding voluntary sterilization):
 - A. Ovarian stimulation and retrieval of oocytes;
 - B. Ovarian tissue retrieval and all of the following:
 - 1. Ovarian tissue is free from malignancy;
 - 2. Insufficient time for oocyte retrieval or member/enrollee is prepubertal;
 - C. Ovarian transposition (oophoropexy).

Note: For those with female reproductive systems \geq age 40 requesting retrieval of their own oocytes, documentation is required noting that the treating provider has evaluated age, infertility risk factors, measure of ovarian reserve, and considers use of the member/enrollee's own oocytes a viable strategy for attempting future conception.

- II. It is the policy of PHW that there is insufficient evidence in the published peer-reviewed literature to support the use of the following procedures for fertility preservation in adults and adolescents with a female reproductive system prior to commencing treatment that is likely to affect fertility:
 - A. Ovarian suppression with gonadotropin releasing hormone (GnRH) agonist or antagonists.
- III. It is the policy of PHW that, when a covered benefit under the member's/enrollee's benefit plan contract, the following procedures are **medically necessary** for adults and adolescents with a male reproductive system prior to commencing medically necessary treatment that is likely to cause infertility (excluding voluntary sterilization):
 - A. Sperm extraction and retrieval procedures.
- IV. It is the policy of PHW that there is insufficient evidence in the published peer-reviewed literature to support the use of the following procedures for fertility preservation in adults and



adolescents with a male reproductive system prior to commencing treatment that is likely to affect fertility:

- A. Testicular suppression with GnRH agonist or antagonists;
- B. Reimplantation or grafting of human testicular tissue.

Background

An estimated 4.4% of all new cancer cases occur among adolescents and young adults between the ages of 15 to 39. Cancer patients are surviving at increasing rates, but successful treatment in younger patients can often be gonadotoxic and lead to late and long-term effects such as infertility. Treatment can affect fertility by causing damage to immature eggs and reproductive organs and affecting the body's hormones. Fertility preservation is an essential part of the management of adolescents and young adults who are at risk for infertility due to cancer treatments.¹¹

Gonadotoxic treatments include chemotherapy, radiation, and surgical resection (for treatment of disease or gender affirmation surgery). Additionally, chemotherapy can be used for noncancerous conditions such as autoimmune diseases, like systemic lupus erythematosus (SLE), and hematological diseases. Prompt counseling regarding available options for fertility preservation for iatrogenic infertility should be provided to patients prior to undergoing any gonadotoxic treatments.¹⁰

American Society for Reproductive Medicine (ASRM)^{10*}

The 2019 ASRM committee opinion for Fertility Preservation in Patients Undergoing Gonadotoxic Therapy or Gonadectomy affirmed that ovarian tissue cryopreservation is no longer considered experimental for prepubertal girls and for those who cannot delay cancer treatment to undergo ovarian stimulation and oocyte retrieval. The committee states, "data on the efficacy, safety, and reproductive outcomes after ovarian tissue cryopreservation are still limited. Given the current body of literature, ovarian tissue cryopreservation should be considered an established medical procedure with limited effectiveness that should be offered to carefully selected patients."

The guideline states that the use of gonadotropin releasing hormone (GnRH) analogs for ovarian protection during chemotherapy remains controversial: "further studies are required to establish the efficacy of this treatment and to determine which patients are the best candidates for its use." Furthermore, GnRH analog therapy for fertility preservation in males has failed to demonstrate effectiveness.

American Society of Clinical Oncology (ASCO)^{5*}

The ASCO recommends discussing fertility preservation with all patients of reproductive age (and with parents or guardians of children and adolescents) if infertility is a potential risk of therapy, as early as possible, before treatment starts.

For those with a male reproductive system who express an interest in fertility preservation, sperm cryopreservation is the only established fertility preservation method. ASCO notes that in these patients, hormonal therapy has not shown to be successful in preserving fertility. Per



ASCO, other methods, including testicular tissue cryopreservation for the purpose of future reimplantation or grafting of human testicular tissue, are experimental.

For those with a female reproductive system who express an interest in fertility preservation, both embryo and oocyte cryopreservation are established fertility preservation methods. The ASCO notes that evidence for ovarian tissue cryopreservation for the purpose of future transplantation remains insufficient. They note also, there is insufficient evidence regarding the effectiveness of ovarian suppression with GnRH agonist or antagonists to preserve fertility.

National Comprehensive Cancer Network (NCCN)^{8*}

NCCN guidelines on Adolescent and Young Adult Oncology note that oocyte or embryo cryopreservation is recommended for those that can delay cancer therapy for approximately three weeks. Ovarian tissue cryopreservation is a promising strategy for fertility preservation when there is insufficient time for oocyte or embryo cryopreservation and/or the patient is prepubertal. Hormonal stimulation is not required with this technique, therefore there is no delay in the initiation of treatment. This procedure is not appropriate for certain patients, including carriers of BRCA mutations due to the increased risk of ovarian cancer and those with cancer if potential exists for reintroduction of malignant cells with grafting. While ovarian tissue cryopreservation is still considered investigational at some institutions, it may be discussed as an option for fertility preservation.

Some data suggests menstrual suppression with GnRH agonists may protect ovarian function. However, evidence that menstrual suppression with GnRH agonists provides adequate protection of the ovaries is controversial, so this procedure is not currently considered a form of fertility preservation.

American College of Obstetricians and Gynecologists $(ACOG)^{6*}$

ACOG's Gynecologic Issues in Children and Adolescent Cancer Patients and Survivors committee opinion states that "cryopreservation of oocytes or embryos may be offered before cancer treatments if there is adequate time and a safe method for ovarian stimulation. Ovarian tissue extraction and cryopreservation have been shown to have some success with posttreatment autotransplantation after chemotherapy."

For young individuals with a female reproductive system who have completed sexual development, GnRH agonists and antagonists such as leuprolide acetate, have been used to induce ovarian quiescence to preserve ovarian function and fertility after cytotoxic treatment. Leuprolide acetate is not recommended prior to puberty. There still is no conclusive evidence that demonstrates efficacy of GnRH agonists and antagonists, and studies are primarily observational regarding their effectiveness in fertility preservation.

Coding Implications

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informational purposes only. Inclusion or exclusion of any codes does not guarantee coverage. Providers should reference the most up-to-date sources of professional coding guidance prior to the submission of claims for reimbursement of covered services.

CPT ®	Description			
Codes				
00840	Anesthesia for intraperitoneal procedures in lower abdomen including			
	laparoscopy; not otherwise specified			
00922	Anesthesia for procedures on male genitalia (including open urethral procedures);			
	seminal vesicles			
53899	Unlisted procedure, urinary system			
55870	Electroejaculation			
55899	Unlisted procedure, male genital system			
58825	Transposition, ovary(s)			
58970	Follicle Puncture for oocyte retrieval, any method			
76856	Ultrasound, pelvic (nonobstetric), real time with image documentation; complete			
76948	Ultrasonic guidance for aspiration of ova, imaging supervision and interpretation			
82670	Estradiol; total			
83001	Gonadotropin; follicle stimulating hormone (FSH)			
83002	Gonadotropin; luteinizing hormone (LH)			
84144	Progesterone			
84702	Gonadotropin; chorionic (hCG); quantitative			
89254	Oocyte identification from follicular fluid			
89320	Semen analysis; volume, count motility and differential			
99000	Handling and/or conveyance of specimen for transfer from office to a laboratory			
99001	Handling and/or conveyance of specimen for transfer from the patient in other than			
	an office to a laboratory (distance may be indicated)			
99070	Supplies and materials (except spectacles), provided by the physician or other			
	qualified health care professional over and above those usually included with the			
	office visit or other services rendered (list drugs, trays, supplies, or materials			
	provided)			
99078	Physician or other qualified health care professional qualified by education,			
	training, licensure/regulation (when applicable) educational services in a group			
	setting (eg, prenatal, obesity, or diabetic instructions)			
99199	Unlisted special service, procedure or report			

HCPCS Codes	Description
S4028	Microsurgical epididymal sperm aspiration (MESA)



CPT	Description
Codes	
that do	
not	
support	
medical	
necessity	
CPT®	
Codes	
53899	Unlisted procedure, urinary system
55899	Unlisted procedure, male genital system
89398	Unlisted reproductive medicine laboratory procedure

Reviews, Revisions, and Approvals	Date	Approval Date
Policy developed.	10/18	
Annual review completed. Codes reviewed. References reviewed	12/19	07/02/2021
and updated. Specialty review completed.		
Removed CPT 0375T – code deleted 1/1/20; Replaced 'members'	06/2021	09/01/2021
with "members/enrollees" in all instances; Revised description of		*Does not
CPT-82670. CPT-0058T deleted in 2021.		require
"Experimental/investigational" verbiage replaced with descriptive		OLTL_OM
language in policy statement III and IV. References reviewed and		AP review.
updated.		
Annual review. Replaced all instances of female and male with	05/2023	
descriptive, gender-neutral verbiage. Added code 89398 to table of		
CPT codes considered investigational. References reviewed and		
updated. In description, removed "male and female" from		
introductory sentence about medical causes of impaired fertility.		
Specified in sections I. and III. that the treatment causing risk to		
fertility was medically necessary. Removed "embryo		
cryopreservation", "cryopreservation of mature oocytes", "conservative gynecologic surgery (radical trachelectomy and		
ovarian cystectomy)" and "radiation (gonadal) shielding" from		
section I. Added "ovarian stimulation and retrieval of oocytes" and		
"ovarian tissue retrieval" to section I. Included "Note: For those with		
female reproductive systems > age 40" to criteria and background		
sections. Removed "cryopreservation of immature oocytes" and		
ovarian tissue cryopreservation and transplantation procedures" from		
section II. Added "sperm extraction procedures and retrieval		
procedures" to section III and removed "cryopreservation of sperm"		
and radiation (gonadal) shielding". Removed "testicular tissue or		
spermatogonial cryopreservation" from section IV. Criteria section		
reformatted for organizational purposes. Background updated. Added		
CPT codes 00922, 53899, 55899, and 55870. Removed CPT codes		
57531, 77334, 89250, 89251, 89258, 89259, 89268, 89272, 89280,		



Reviews, Revisions, and Approvals	Date	Approval Date
89281, 89337, 89352, 89353. Added HCPCS codes S4028 and removed HCPCS codes S4030 and S4031. Added CPT code 53899 and 55899 and removed 89335 from the "does not support" table. References reviewed and updated. Internal specialist review.		

References

- 1. Practice Committee of American Society for Reproductive Medicine. Ovarian tissue cryopreservation: a committee opinion. Fertil Steril. 2014;101(5):1237 to1243. doi:10.1016/j.fertnstert.2014.02.052
- 2. Shah JS, Guerra R, Bodurka DC, Sun CC, Chisholm GB, Woodard TL. Factors influencing fertility-sparing treatment for gynecologic malignancies: A survey of Society of Gynecologic Oncology members. Gynecol Oncol. 2017;147(3):497 to502. doi:10.1016/j.ygyno.2017.09.019
- 3. Health Technology Assessment. Ovarian tissue cryopreservation for preservation of fertility in patients undergoing gonadotoxic cancer treatment. Hayes. www.hayesinc.com. Published October 1, 2019 (annual review December 8, 2022). Accessed April 18, 2023.
- 4. Pacheco F, Oktay K. Current Success and Efficiency of Autologous Ovarian Transplantation: A Meta-Analysis. Reprod Sci. 2017;24(8):1111 to1120. doi:10.1177/1933719117702251
- 5. Oktay K, Harvey BE, Partridge AH, et al. Fertility Preservation in Patients with Cancer: ASCO Clinical Practice Guideline Update. J Clin Oncol. 2018;36(19):1994 to2001. doi:10.1200/JCO.2018.78.1914
- 6. The American College of Obstetricians and Gynecologists. Gynecologic Issues in Children and Adolescent Cancer Patients and Survivors No.747. www.acog.org. Published July 2018 (reaffirmed 2021). Accessed April 17, 2023.
- 7. Sonmezer M, Oktay K. Fertility and reproductive hormone preservation: Overview of care prior to gonadotoxic therapy or surgery. UpToDate. www.uptodate.com. Published July 8, 2022. . Accessed April 17, 2023.
- 8. National Comprehensive Cancer Network. Adolescent and Young Adult (AYA) Oncology (Version 3.2023). https://www.nccn.org/guidelines/category 4. Accessed April 17, 2023.
- 9. Oktay K, Sonmezer M. Fertility preservation: Cryopreservation options. UpToDate. www.uptodate.com. Published February 13, 2023. Accessed April 17, 2023.
- 10. Practice Committee of the American Society for Reproductive Medicine. Electronic address: asrm@asrm.org. Fertility preservation in patients undergoing gonadotoxic therapy or gonadectomy: a committee opinion. Fertil Steril. 2019;112(6):1022 to 1033. doi:10.1016/j.fertnstert.2019.09.013
- 11. National Cancer Institute. Adolescents and Young Adults with Cancer. https://www.cancer.gov/types/aya. Published April 24, 2023. Accessed May 5, 2023.