

Comprehensive Fertility Care and Preservation for Male Cancer Survivors

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Disclosure Statement

- We do not have any conflict of interest, nor will we be discussing any offlabel product use.
- This class has no commercial support or sponsorship, nor is it cosponsored

Survivorship and Fertility

In a survey questioning the importance of fertility preservation in women of childbearing age recently diagnosed with cancer.



77.6% (66 of 85) reported that the possibility of preserving their fertility was instrumental to coping before and after treatment



Treves R et. al., *Future Oncol (*2014) K. Gard. Photograph of UW Male Fertility Lab. (2021)

Testicular tissue serves a role in both hormone production and reproduction

Ovary

- Reproduction
 - Females born with lifetime oocytes
- Hormones
 - Progesterone
 - Estrogen (estradiol, estrone, and estriol)

Testes

- Reproduction
 - Ongoing production of sperm after puberty
- Hormone
 - Testosterone







Spermatogenesis



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Breuss et al. Trends in Genetics (2021)

Risk Quantification and Gonadal Failure



Gonadotoxic Therapy





CI1

Nitromin

Cyclophosphamide



CISPLATIN & CARBOPLATIN



SURGERY



ALKYLATING CHEMOTHERAPY



Male Risk Stratification

		Minimally Increased Risk	Significantly Increased risk	High level of Increased risk
Alkylators CED* gm/m2		CED < 4		CED ≥ 4
Hematopoietic Stem Cell Transplant				Alkylator +/-Total body irradiation Myeloablative and Reduced intensity regimens
Heavy Metal mg/m2		Cisplatin Carboplatin	Cisplatin > 500	
Radiation Exposure	Testicular	0.2-0.6Gy	0.7-3.9 Gy	≥ 4.0 Gy
	Hypothalamus	26-29.99	> 30-39.9 Gy	> 40 Gy
Surgery			RPLND [^]	





Cyclophosphamide Equivalent Dose

Cyclophosphamide Equivalent Dose Calculator								
Dose			Equivalent dose					
Drug	(mg/m2)	Multiplier	(mg/m2)					
Cyclophosphamide		1	0					
Ifosfamide		0.244	0					
Procarbazine		0.857	0					
Chlorambucil		14.286	0					
BCNU		15	0					
CCNU		16	0					
Melphalan		40	0					
Thio-TEPA		50	0					
Nitrogen Mustard		100	0					
Busulfan		8.823	0					

Total CED 0



https://fertilitypreservationpittsburgh.org/fertility-resources/fertility-risk-calculator/ Green DM, et al. Pediatric Blood & Cancer (2014)

Male Infertility Risk: Principles

- Long term outcomes: mostly binary (fertile vs not fertile)
- Risks do not vary greatly with age
- Temporary azoospermia is of variable length.





Ntemou et al. International Journal of Molecular Science (2019)

Options for Males

Before Treatment Options	Potential Use	Dis- advantages	Status
Sperm Freezing (Semen/Testis)	IUI, IVF	Limited Sperm	Standard
Prepub Testicular Tissue Freezing	IVF	Few sperm Surgery	Experimental

After Treatment Options	Fresh Sperm	Frozen Sperm	Status
Diagnostic Semen Analysis	\checkmark		Standard
Natural Conception	\checkmark		Standard
IntraUterine Insemination IUI	\checkmark	\checkmark	Standard
In Vitro Fertilization IVF	\checkmark	\checkmark	Standard
Donor sperm IUI or IVF		\checkmark	Standard
Adoption			Standard

Sperm cryopreservation





- Must be Tanner Stage 3
- Masturbated previously
- No lubricants or saliva
- Sperm must be collected prior to any chemotherapy



Testicular Tissue Cryopreservation



Offered by experimental protocol only No human live births to date.



Ntemou et al. International Journal of Molecular Science (2019)

Comprehensive Fertility Preservation at Seattle Children's Hospital

MALE

Prepubertal

• Testicular Tissue Cryopreservation (Experimental - Protocol at SCH)

Pubertal

- Sperm Cryopreservation
- Testicular Sperm Extraction/Aspiration (TESE/TESA)
- Testicular Tissue Cryopreservation (Experimental - Protocol at SCH)

Comprehensive evaluation and evaluation of all patients facilitated by Oncofertility team including oncology, general surgery, urology, and reproductive endocrinology.



Resources

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Seattle Children's Fertility Preservation Team FertilityPreservationRN@seattlechildrens.org Kristin.gard@seattlechildrens.org UW Male Fertility Lab www.malefertilitylab.com Seattle Reproductive Medicine

www.seattlefertility.com





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