NCCN Guidelines Version 3.2025 Hereditary Cancer Testing Criteria

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Discussion

TESTING CRITERIA FOR PROSTATE CANCER SUSCEPTIBILITY GENES (Genes such as *ATM*, *BRCA1*, *BRCA2*, *CHEK2*, *HOXB13*, bb and *TP53*) (GENE-A)a,cc,dd

Testing is clinically indicated in the following scenarios:

- See General Tumor Criteria on CRIT-1.
- Personal history of prostate cancer at any age with:
- ▶ Metastatic^p (Stage IVB) or node-positive (Stage IVA) prostate cancer
- ▶ Very high-risk or high-risk disease (See Initial Risk Stratification and Staging Workup [PROS-2] NCCN Guidelines for Prostate Cancer)
- ▶ Ancestry: Ashkenazi Jewish
- **▶** Family history
 - **♦ ≥1 close blood relative** with ANY:
 - breast cancer at age ≤50 y
 - male breast cancer
 - ovarian cancer
 - pancreatic cancer
 - metastatic,^p node positive, or very high-risk or high-risk prostate cancer
 - ♦ ≥3 close blood relatives^o with prostate cancer (any grade) and/or breast cancer on the same side of the family including the patient with prostate cancer
 - ♦ For additional family history considerations, see NCCN Guidelines for Genetic/Familial High-Risk Assessment: Colorectal, Endometrial, and Gastric
- Family history criteria: unaffected; or affected but does not meet criteria above
 - ♦ Individual with a first-degree blood relative meeting any of the criteria listed above (except unaffected individuals whose relatives meet criteria only for systemic therapy decision-making)^q

Testing may be considered in the following scenario:

- Personal history of prostate cancer ≤55 yee not meeting any of the above criteria
- Personal history of prostate cancer with intermediate-risk prostate cancer with intraductal/cribriform histology (see Initial Risk Stratification and Staging Workup in NCCN Guidelines for Prostate Cancer) at any age

Refer for genetic testing in the following scenario:

- Biochemical recurrence
- ▶ For this heterogeneous disease state, refer for genetic testing if: 1) Very high-risk or high-risk disease at diagnosis (see above), 2) family history of cancer criteria met, or if (and at time of) progression to metastatic disease

Criteria → GENE-1 met If criteria for other If testing hereditary criteria syndromes not met. not met. consider then testing cancer criteria screening for other as per hereditary NCCN syndromes Screening Guidelines

Footnotes on CRIT-6A