# **Enhancing Breast Cancer Risk Assessment in a Community Imaging Center to** Identify High-Risk Patients and Guide Screening and Management

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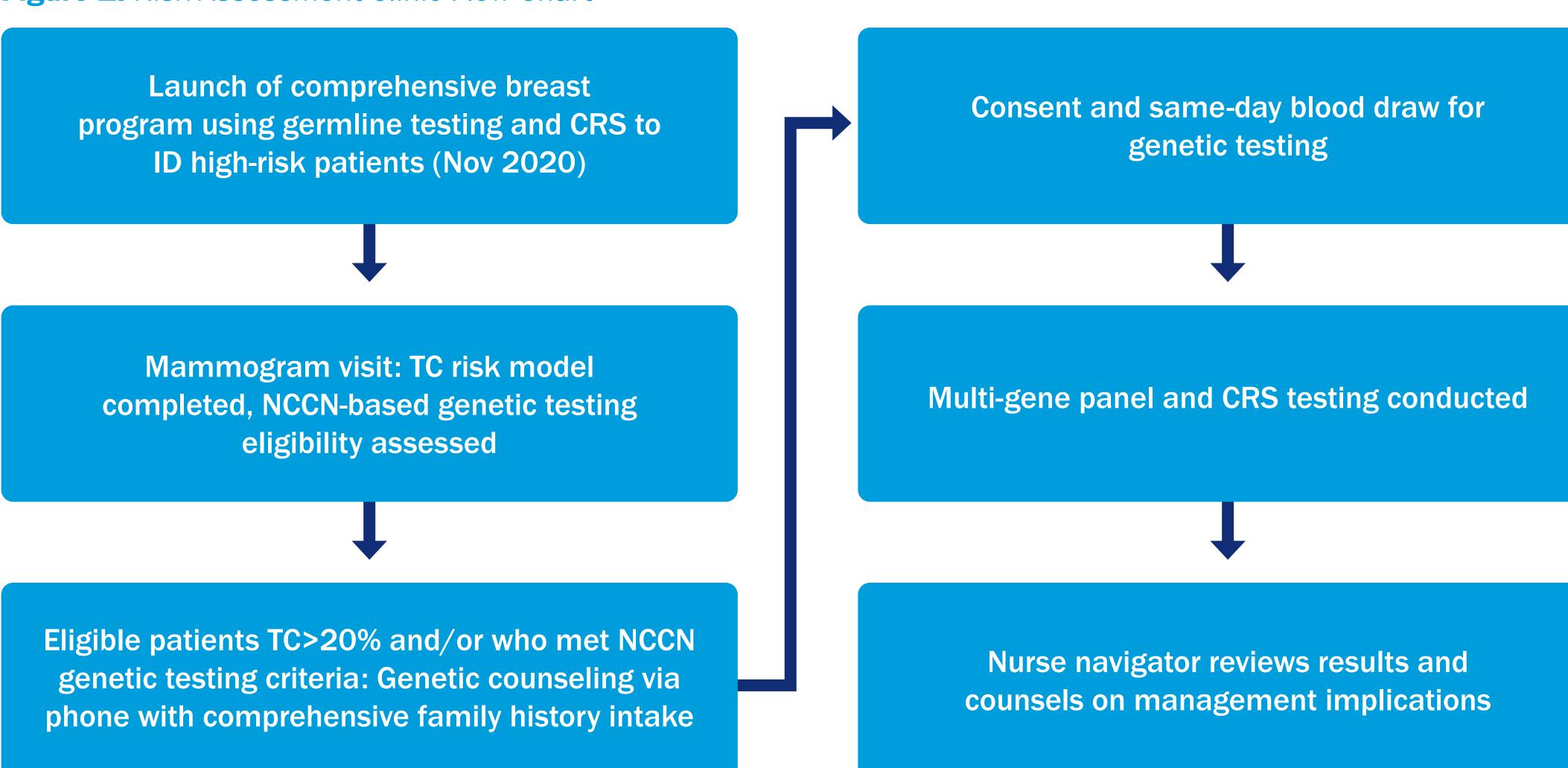
# Background

- To assess the effectiveness of Tyrer-Cuzick version 8 (TC) alone versus a combined approach—including family history assessment, germline testing, and TC augmented with a breast cancer multiple ancestry polygenic risk score (MA-PRS)—in identifying patients at elevated risk for breast and other cancers at imaging centers. TC+MA-PRS = Combined Risk Score (CRS).
- Here we compared approaches for identifying high-risk patients to improve personalized screening and management.

# Methods

- In November 2020, Singing River Health System launched a comprehensive breast program using the TC risk model and germline testing with CRS to identify high-risk patients. (Figure 1).
- During mammogram visits, patients completed the TC risk model and were assessed for genetic testing eligibility based on NCCN guidelines using a cancer history questionnaire (**Figure 2**).
- Eligible patients received a comprehensive family history intake and education by telephone from a genetic counselor and, if consented, had blood drawn for genetic testing the same day (Figure 3).
- The testing included a laboratory's multi-gene panel and CRS to determine BC risk and hereditary cancer syndromes (Figure 4).
- Post-testing, a nurse navigator reviewed the results and counseled patients on appropriate next steps.
- We evaluated results to identify guideline-recommended management changes and to determine if highrisk patients would have been missed without germline and CRS.

#### Figure 1. Risk Assessment Clinic Flow Chart



## • From November 2020 through December 2023, a total of 2,434 patients completed TC and if they met the criteria for genetic testing at the time of their mammogram, they had multi-gene panel testing and CRS.

• Within this cohort, 677 patients had a TC < 20% lifetime BC risk, yet were candidates for medical management change based on their genetic testing, CRS, and/or comprehensive family history results.

### Figure 2. Family History Questionnaire

| (The following relatives should be considered: Parents, siblings, half-siblings, children, grandparents, grandchildren, aunts, uncles, nieces, nephews, on both sides of the family)  Yes or No  | Updated May 2025  Breast Cancer diagnosed at age 65 or younger?  Triple Negative Breast Cancer at any age?  Bilateral (two breast cancers) at any age?  Metastatic (stage 4) Breast Cancer?  Ovarian Cancer at any age?  Pancreatic Cancer at any age?  Colon/Colorectal Cancer at any age?  Endometrial Cancer at any age?  Endometrial Cancer at any age?  Have YOU been tested for hereditary risk of cancer? Yes No  (Genetic Testing/ e.g. BRCA or Lynch Syndrome) If so, when:  Do you have a Family History of any of the following?  (The following relatives should be considered: Parents, siblings, half-sibling grandchildren, aunts, uncles, nieces, nephews, on both sides of the family) | YesNoYesNoYesNoYesNoYesNoYesNoYesNoYesNoYesNoYesNoYesNoYesNoYesNoYesNoYesNoYesNoYesNoResult:  | <i>lparents,</i>       |
|--|---|---|------------------------|
| Breast Cancer diagnosed at age 65 or younger?  Triple Negative Breast Cancer at any age?  Ves No  Bilateral (two breast cancers) at any age?  Ves No  Metastatic (stage 4) Breast Cancer?  Ves No  Metastatic (stage 4) Breast Cancer?  Ves No  Pancreatic Cancer at any age?  Ves No  Colon/Colorectal Cancer at any age?  Ves No  Colon/Colorectal Cancer at any age?  Ves No  10 or more colon polyps in your lifetime?  Wes No  10 or more colon polyps in your lifetime?  Wes No  Have YOU been tested for hereditary risk of cancer? Yes No  Colon/Colorectal Cancer at any age?  Fish of No  Have YOU been tested for hereditary risk of cancer? Yes No  How You have a Family History of any of the following:  Wes or No  Which Relative?  Which Relative?  Which Relative?  Breast Cancer at any age.  Yes No  Maternal (Mother) or Paternal (Father)  Paternal (Father)  Paternal (Father)  Age at Diagnosis?  Paternal (Father)  Paternal (Father)  Paternal (Father)  Age at Diagnosis?  Paternal (Father)  Paternal (Father)  Age at Diagnosis?  Paternal (Father)  Paternal (Father | Breast Cancer diagnosed at age 65 or younger?  Triple Negative Breast Cancer at any age?  Bilateral (two breast cancers) at any age?  Metastatic (stage 4) Breast Cancer?  Ovarian Cancer at any age?  Pancreatic Cancer at any age?  Colon/Colorectal Cancer at any age?  Endometrial Cancer at any age?  10 or more colon polyps in your lifetime?  Have YOU been tested for hereditary risk of cancer? Yes No (Genetic Testing/ e.g. BRCA or Lynch Syndrome) If so, when:  Do you have a Family History of any of the following?  (The following relatives should be considered: Parents, siblings, half-sibling grandchildren, aunts, uncles, nieces, nephews, on both sides of the family)         | YesNoYesNoYesNoYesNoYesNoYesNoYesNoYesNoYesNoYesNoYesNoYesNoYesNoYesNoYesNoYesNoYesNoResult:  | <i>lparents,</i>       |
| Breast Cancer diagnosed at age 65 or younger?  | Triple Negative Breast Cancer at any age?  Bilateral (two breast cancers) at any age?  Metastatic (stage 4) Breast Cancer?  Ovarian Cancer at any age?  Pancreatic Cancer at any age?  Colon/Colorectal Cancer at any age?  Endometrial Cancer at any age?  10 or more colon polyps in your lifetime?  Have YOU been tested for hereditary risk of cancer? Yes No (Genetic Testing/ e.g. BRCA or Lynch Syndrome) If so, when:  Do you have a Family History of any of the following? (The following relatives should be considered: Parents, siblings, half-sibling grandchildren, aunts, uncles, nieces, nephews, on both sides of the family)   | Yes No Result:  ings, children, grand Maternal (Mother) or Paternal (Father) | <i>lparents,</i>       |
| Bilateral (two breast cancers) at any age?   | Bilateral (two breast cancers) at any age?  Metastatic (stage 4) Breast Cancer?  Ovarian Cancer at any age?  Pancreatic Cancer at any age?  Colon/Colorectal Cancer at any age?  Endometrial Cancer at any age?  10 or more colon polyps in your lifetime?  Have YOU been tested for hereditary risk of cancer? Yes No  (Genetic Testing/ e.g. BRCA or Lynch Syndrome) If so, when:  Do you have a Family History of any of the following?  (The following relatives should be considered: Parents, siblings, half-sibling grandchildren, aunts, uncles, nieces, nephews, on both sides of the family)  | Yes No Result:  ings, children, grand Maternal (Mother) or Paternal (Father)               | -                      |
| Metastatic (stage 4) Breast Cancer?  Ovarian Cancer at any age?  Pes No  Ovarian Cancer at any age?  Yes No  Colon/Colorectal Cancer at any age?  Yes No  Endometrial Cancer at any age?  Yes No  To or more colon polyps in your lifetime?  Have YOU been tested for hereditary risk of cancer? Yes No  (Genetic Testing/ e.g. BRCA or Lynch Syndrome)  Have YOU been tested for hereditary risk of cancer? Yes No  (Genetic Testing/ e.g. BRCA or Lynch Syndrome)  Have YOU been tested for hereditary risk of cancer? Yes No  (Genetic Testing/ e.g. BRCA or Lynch Syndrome)  Have YOU been tested for hereditary risk of cancer? Yes No  (Genetic Testing/ e.g. BRCA or Lynch Syndrome)  Have YOU been tested for hereditary risk of cancer? Yes No  Have YOU been tested for hereditary risk of cancer? Yes No  Have YOU been tested for hereditary risk of cancer? Yes No  Have YOU been tested for hereditary risk of cancer? Yes No  Have YOU been tested for hereditary risk of cancer? Yes No  Have YOU been tested for hereditary risk of cancer? Yes No  Have YOU been tested for hereditary risk of cancer? Yes No  Have YOU been tested for hereditary risk of cancer? Yes No  Which Relatives?  Maternal (Mother) or Paternal (Mother) or Paternal (Mother) or Paternal (Father)  Age at Diagnosis?  Paternal (Hother) or Maternal (Mother) or Paternal (Father)  Age at Diagnosis?  Age at Diagnos | Metastatic (stage 4) Breast Cancer?  Ovarian Cancer at any age?  Pancreatic Cancer at any age?  Colon/Colorectal Cancer at any age?  Endometrial Cancer at any age?  In or more colon polyps in your lifetime?  Have YOU been tested for hereditary risk of cancer? Yes No  (Genetic Testing/ e.g. BRCA or Lynch Syndrome)  If so, when:  O you have a Family History of any of the following?  The following relatives should be considered: Parents, siblings, half-sibling grandchildren, aunts, uncles, nieces, nephews, on both sides of the family)   | Yes No Result:  ings, children, grand Maternal (Mother) or Paternal (Father)                      | -                      |
| Ovarian Cancer at any age? Pancreatic Cancer at any age? Pancreatic Cancer at any age? Pancreatic Cancer at any age? Yes No Colon/Colorectal Cancer at any age? Yes No Endometrial Cancer at any age? Yes No 10 or more colon polyps in your lifetime? Yes No Have YOU been tested for hereditary risk of cancer? Yes No (Genetic Testing) e.g. BRCA or Lynch Syndrome) If so, when:   | Ovarian Cancer at any age?  Pancreatic Cancer at any age?  Colon/Colorectal Cancer at any age?  Endometrial Cancer at any age?  10 or more colon polyps in your lifetime?  Have YOU been tested for hereditary risk of cancer? Yes No  (Genetic Testing/ e.g. BRCA or Lynch Syndrome) If so, when:  O you have a Family History of any of the following?  The following relatives should be considered: Parents, siblings, half-sibling grandchildren, aunts, uncles, nieces, nephews, on both sides of the family)   | Yes No Yes No Yes No Yes No Yes No Yes No Result:  ings, children, grand Maternal (Mother) or Paternal (Father)                             | -                      |
| Pancreatic Cancer at any age?  Colon/Colorectal Cancer at any age?  Yes No  Endometrial Cancer at any age?  Yes No  10 or more colon polyps in your lifetime?  Have YOU been tested for hereditary risk of cancer? Yes No  (Genetic Testing/ e.g. BRCA or Lynch Syndrome) If so, when: Result:   | Pancreatic Cancer at any age?  Colon/Colorectal Cancer at any age?  Endometrial Cancer at any age?  10 or more colon polyps in your lifetime?  Have YOU been tested for hereditary risk of cancer? Yes No  (Genetic Testing/ e.g. BRCA or Lynch Syndrome) If so, when:  O you have a Family History of any of the following?  The following relatives should be considered: Parents, siblings, half-siblings, the following relatives, nieces, nephews, on both sides of the family)  | Yes No Yes No Yes No Yes No Yes No Result: ings, children, grand Maternal (Mother) or Paternal (Father)                                     | -                      |
| Colon/Colorectal Cancer at any age?  | Colon/Colorectal Cancer at any age?  Endometrial Cancer at any age?  10 or more colon polyps in your lifetime?  Have YOU been tested for hereditary risk of cancer? Yes No  (Genetic Testing/ e.g. BRCA or Lynch Syndrome) If so, when:  Oo you have a Family History of any of the following?  The following relatives should be considered: Parents, siblings, half-sibling trandchildren, aunts, uncles, nieces, nephews, on both sides of the family)   | Yes No Yes No Yes No Result: Maternal (Mother) or Paternal (Father)   | -                      |
| Endometrial Cancer at any age?  10 or more colon polyps in your lifetime?  Have YOU been tested for hereditary risk of cancer? Yes No  | Endometrial Cancer at any age?  10 or more colon polyps in your lifetime?  Have YOU been tested for hereditary risk of cancer? Yes No  (Genetic Testing/ e.g. BRCA or Lynch Syndrome) If so, when:  Oo you have a Family History of any of the following?  The following relatives should be considered: Parents, siblings, half-sibling trandchildren, aunts, uncles, nieces, nephews, on both sides of the family)  | Yes No Yes No Result: Maternal (Mother) or Paternal (Father)  | -                      |
| Have YOU been tested for hereditary risk of cancer? YesNo  | 10 or more colon polyps in your lifetime?  Have YOU been tested for hereditary risk of cancer? Yes No (Genetic Testing/ e.g. BRCA or Lynch Syndrome) If so, when:  Oo you have a Family History of any of the following?  The following relatives should be considered: Parents, siblings, half-siblings, half-siblings, half-siblings, aunts, uncles, nieces, nephews, on both sides of the family)  | Yes No  Result:  Maternal (Mother) or Paternal (Father)   | -                      |
| Have YOU been tested for hereditary risk of cancer? Yes  | Have <b>YOU</b> been tested for <b>hereditary risk of cancer</b> ? Yes No (Genetic Testing/ e.g. BRCA or Lynch Syndrome) If so, when: Do you have a <b>Family History</b> of any of the following?  The following relatives should be considered: Parents, siblings, half-siblings, and children, aunts, uncles, nieces, nephews, on both sides of the family)  | Result: ings, children, grand Maternal (Mother) or Paternal (Father)  | -                      |
| Genetic Testing/ e.g. BRCA or Lynch Syndrome    If so, when: Result:   | (Genetic Testing/e.g. BRCA or Lynch Syndrome) If so, when:  Do you have a <b>Family History</b> of any of the following?  The following relatives should be considered: Parents, siblings, half-siblings, aunts, uncles, nieces, nephews, on both sides of the family)  | ings, children, grand Maternal (Mother) or Paternal (Father)  | -                      |
| Do you have a Family History of any of the following?  The following relatives should be considered: Parents, siblings, half-siblings, children, grandparents, grandchildren, aunts, uncles, nieces, nephews, on both sides of the family)  Yes or No   Which Relative?   Maternal (Mother) or Paternal (Father)    Paternal (Father)   Paternal (Father)    Breast cancer age 49 or younger   Yes   No     M   P    2 Breast Cancers in one relative at any age   Yes   No     M   P    3 breast or prostate cancers in   Yes   No     M   P    7 Pancreatic cancer at any age   Yes   No     M   P    Pancreatic cancer at any age   Yes   No     M   P    Pancreatic cancer at any age   Yes   No     M   P    Pancreatic cancer at any age   Yes   No     M   P    Pancreatic cancer at any age   Yes   No     M   P    Pancreatic cancer at any age   Yes   No     M   P    Pancreatic cancer at any age   Yes   No     M   P    Pancreatic cancer at any age   Yes   No     M   P    Pancreatic cancer at any age   Yes   No     M   P    Pancreatic cancer at any age   Yes   No     M   P    Pancreatic cancer at any age   Yes   No     M   P    Pancreatic cancer at any age   Yes   No     M   P    Pancreatic cancer at any age   Yes   No     M   P    Alabel Breast cancer at any age   Yes   No     M   P    Pancreatic cancer at any age   Yes   No     M   P    Pancreatic cancer at any age   Yes   No     M   P    Pancreatic cancer at any age   Yes   No     M   P    Pancreatic cancer at any age   Yes   No     M   P    Pancreatic cancer at any age   Yes   No     M   P    Pancreatic cancer at any age   Yes   No     M   P    Pancreatic cancer at any age   Yes   No     M   P    Pancreatic cancer at any age   Yes   No     M   P    Pancreatic cancer at any age   Yes   No     M   P    Pancreatic cancer at any age   Yes   No     M   P    Pancreatic cancer at any age   Yes   No     M   P    Pancreatic cancer at any age   Yes   No     M   P    Pancreatic cancer at any age   Yes   No     M   P    Pancreatic cancer at any age   Yes   No     M   P    Pancreatic cancer at any age | Do you have a <b>Family History</b> of any of the following?<br>(The following relatives should be considered: Parents, siblings, half-siblings, half-siblings)   | ings, children, grand<br>)<br>Maternal (Mother) or<br>Paternal (Father)   | -                      |
| The following relatives should be considered: Parents, siblings, half-siblings, children, grandparents, grandchildren, aunts, uncles, nieces, nephews, on both sides of the family?  Yes or No   | The following relatives should be considered: Parents, siblings, half-siblings, aunts, uncles, nieces, nephews, on both sides of the family)  | Maternal (Mother) or<br>Paternal (Father)   | -                      |
| Breast cancer age 49 or younger  | Yes or No   Which Relative?   1   | Paternal (Father)   | Age at Diagnosis?      |
| Breast cancer age 49 or younger  |   | ` ` `   |                        |
| any age 3 breast or prostate cancers in relatives on the same side of family Ovarian cancer at any age   | Breast cancer age 49 or younger Yes No  |   |                        |
| relatives on the same side of family Ovarian cancer at any age Pancreatic cancer at any age Pancreatic cancer at any age Pancreatic cancer at any age Pancreatic, children, siblings Metastatic prostate cancer at any age Male Breast cancer at any age Male Breast cancer at any age Male Breast cancer at any age Personal Male Personal Male Breast cancer at any Age Male Breast cancer at any age Personal Male Pe | <del></del>   | MP  |                        |
| Pancreatic cancer at any age inYesNo   | ·   | MP  |                        |
| Metastatic prostate cancer at any age  | Ovarian cancer at any age YesNo   | MP  |                        |
| Male Breast cancer at any ageYesNoMP  Colon or endometrial cancer at ageYesNoMP  49 or younger in parents, children or siblingsMP  Ashkenazi Jewish with breastYesNoMP  cancer any age3 relatives on the same side withYesNoMP  Colon, Endometrial or other Lynch cancer**  **Lynch cancers: Colon, Endometrial, Stomach, Ovarian, Pancreatic, Small Bowel, Renal Pelvis, Brain, Sebaceous Carcinoma for the best of my knowledge, I have provided the most accurate answers to the above questions.  **Lynch cancers: Colon, Endometrial, Stomach, Ovarian, Pancreatic, Small Bowel, Renal Pelvis, Brain, Sebaceous Carcinoma for the best of my knowledge, I have provided the most accurate answers to the above questions.  **Lynch cancers: Colon, Endometrial, Stomach, Ovarian, Pancreatic, Small Bowel, Renal Pelvis, Brain, Sebaceous Carcinoma for the best of my knowledge, I have provided the most accurate answers to the above questions.  **Lynch cancers: Colon, Endometrial, Stomach, Ovarian, Pancreatic, Small Bowel, Renal Pelvis, Brain, Sebaceous Carcinoma for the best of my knowledge, I have provided the most accurate answers to the above questions.  **Date:  | parents, children, siblings   | MP  |                        |
| Colon or endometrial cancer at age   | age   |   |                        |
| 49 or younger in parents, children or siblings  Ashkenazi Jewish with breastYesNoMP  |   |   |                        |
| Ashkenazi Jewish with breastYesNo  | 49 or younger in parents, children  | MP  |                        |
| 3 relatives on the same side with Colon, Endometrial or other Lynch cancer**  ***Lynch cancers: Colon, Endometrial, Stomach, Ovarian, Pancreatic, Small Bowel, Renal Pelvis, Brain, Sebaceous Carcinoma To the best of my knowledge, I have provided the most accurate answers to the above questions. Patient's signature:  **COR OFFICE USE ONLY: Did patient meet criteria for Genetic Education?   | Ashkenazi Jewish with breast YesNo  | MP  |                        |
| **Lynch cancers: Colon, Endometrial, Stomach, Ovarian, Pancreatic, Small Bowel, Renal Pelvis, Brain, Sebaceous Carcinoma To the best of my knowledge, I have provided the most accurate answers to the above questions.  Patient's signature:  Date:  OR OFFICE USE ONLY: Did patient meet criteria for Genetic Education?   | Colon, Endometrial or other Lynch   | MP  |                        |
| To the best of my knowledge, I have provided the most accurate answers to the above questions.  Patient's signature:   |   | l Danal Dalvia Duain (  | Coho ao ava Consin ano |
| Did patient meet criteria for Genetic Education?   | Γο the best of my knowledge, I have provided the most accurate answers to the a   | above questions.  |                        |
| If YES, Patient:   ACCEPTED Video  DECLINED Video: Reason  If ACCEPTED, Patient:  SUBMITTED MyRisk  DECLINED Testing: Reason  ACCEPT or  DECLINE genetic testing.  |   |   |                        |
| f ACCEPTED, Patient:   SUBMITTED MyRisk DECLINED Testing: Reason  ACCEPT or DECLINE genetic testing.   | Did patient meet criteria for Genetic Education? □ YES □ NO □ MORE INF  | FORMATION NEEDED  | )                      |
| f ACCEPTED, Patient:   SUBMITTED MyRisk DECLINED Testing: Reason  ACCEPT or DECLINE genetic testing.   | f YES, Patient:   ACCEPTED Video   DECLINED Video: Reason_  |   |                        |
| ACCEPT orDECLINE genetic testing.  |   |   |                        |
|  |   | ···   |                        |
| Patient's signature: Date: Date:   |   | _   |                        |
|  | Patient's signature:  | Date:   |                        |
|  |   |   |                        |

#### Figure 3. Management Changes in Patients with TC < 20%

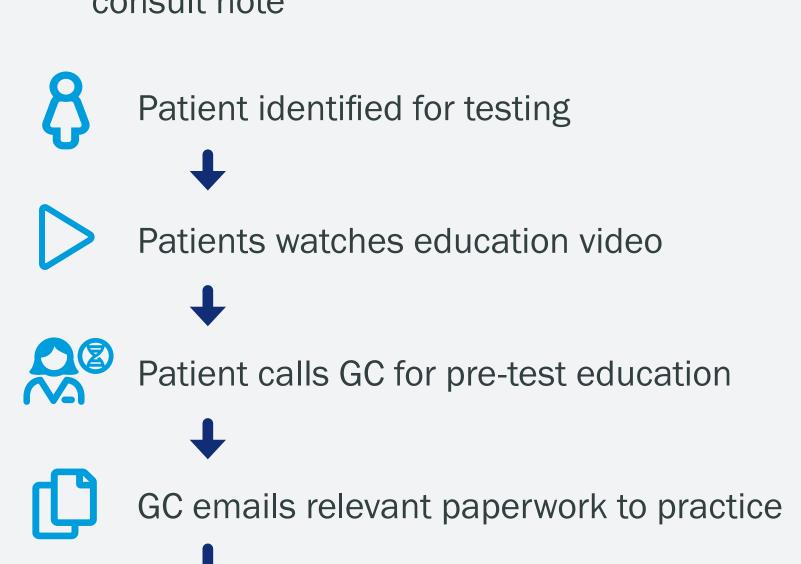
Patients with low risk by TC (<20%) found to be at high-risk by other clinical strategies and eligible for management changes.

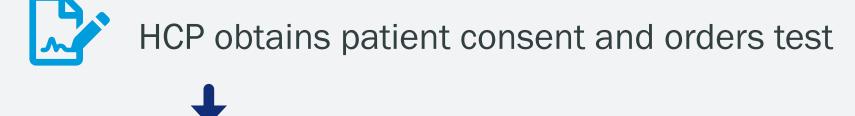
# **Pre-test education**

We offer live, point-of-care pre-test sessions to educate your patients about the test you recommend and answer any questions they may have

#### Myriad's Genetic Counselors will:

- Review patient's family history
- Educate the patient about genetic testing, the recommended test, and the results' impact on medical management
- Email clinic relevant paperwork including a pre-populated test request form and detailed consult note

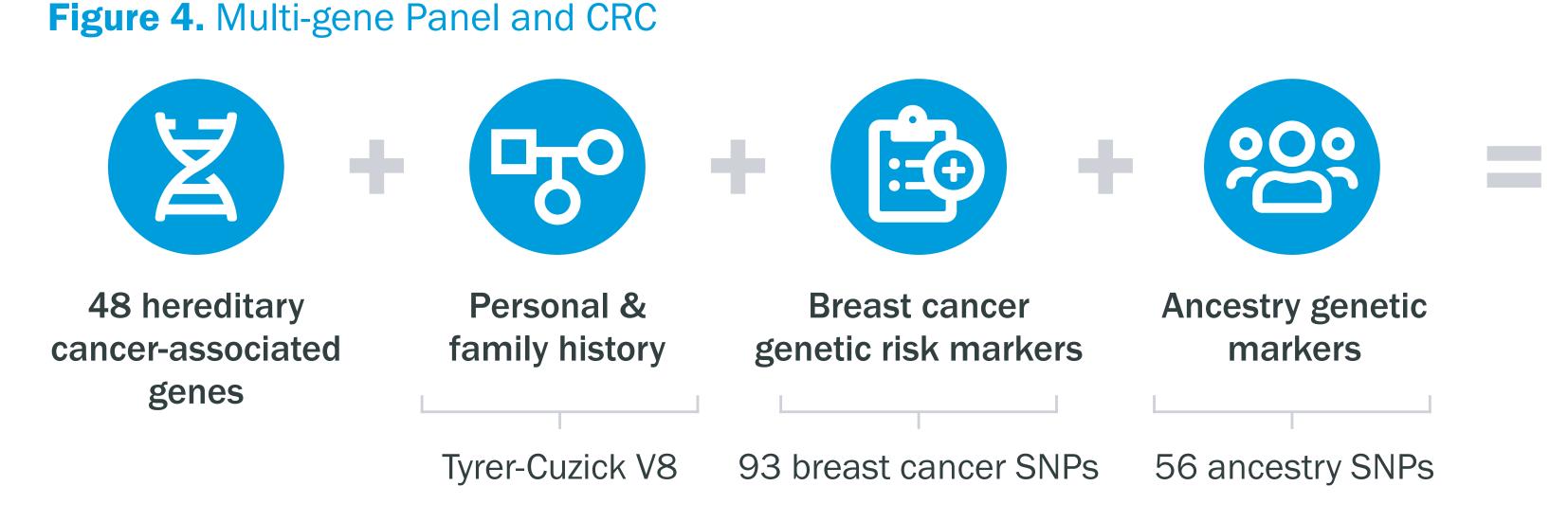




Practice collects patient sample

Practice packages documentation and sample for FedEx delivery to Myriad

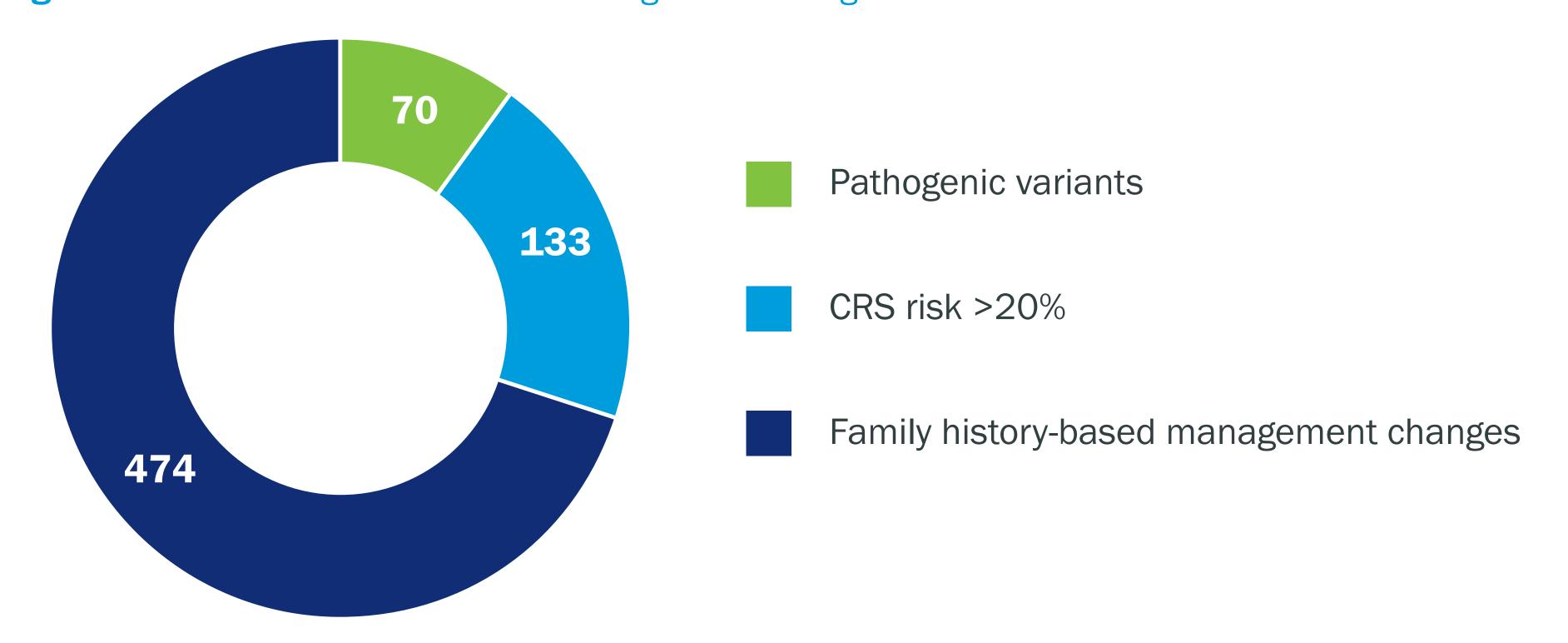
Results



carry pathogenic variants in genes associated with hereditary cancer syndromes.

**Germline genetic** testing with TC8 + MA-**PRS (Multiple Ancestry** Polygenic Risk Score)

#### Figure 5. Patients with TC<20% with Management Changes



• Specifically, 70 (10.34%) patients with TC <20% met the criteria for genetic testing and were found to

• Additionally, 133 (19.63%) patients with a TC result <20% were found to have >20% BC risk by CRS.

based on family history information obtained through patient education intake (Figure 5).

• Furthermore, 474 (70.04%) patients with TC <20% were eligible for modification of management strategy

# Conclusions

- A detailed risk assessment and testing program involving TC, a polygenic risk score model, and germline testing at a community imaging center identified more high-risk individuals and suggested alterations in screening and management compared to using TC alone.
- Without this approach, patients at elevated cancer risk requiring management changes would not have been recognized.