### EAQ152 Available Through ECOG-ACRIN Cancer Research Group

**COMET—Communication and Education in Tumor Profiling:**

A Randomized Study of Pre-disclosure Genetic Education v. Usual Care in Tumor Profiling for Advanced Cancer and a Pilot Study of Remote Genetic Counseling for Participants with Potential Germline Mutations Identified on Tumor Profiling

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**Patient Population**

See Section 3.0 for Complete Eligibility Details

**Step 1 - Primary Intervention Study (RCT):**

- Must be registered to the first screening step (Step 0) for the NCI-MATCH trial (EAY131) OR must be having tumor profiling for advanced cancer at one of the expansion sites listed per protocol
- Must speak English; must have web and e-mail access
- Patients in the MATCH trial must not have received his/her MATCH tumor genetic testing results; non-MATCH expansion site patients must not have received the results from tumor profile genetic testing

**Step 2 - Secondary Genetic Counseling Substudy:**

- Only available to patients from select participating sites (see Section 4.2.3.2)
- Must meet the eligibility criteria above (except for having web/email access and not receiving the tumor genetic testing results); must have had the results from their tumor genetic testing shared with them (non-MATCH expansion sites must have participated in COMET Step 1); must be able to hear by phone
- Must have a potential germline mutation, as determined by the MATCH tumor profiling assay/other clinical lab, and must meet 1 of the following criteria:
  - Tumor contains BRCA1, BRCA2, MLH1, MSH2, TSC1*, TSC2*, VHL*, CDH1, CDKN2A*  
  - Has an APC mutation and NOT colon cancer  
  - Has an APC mutation, colon cancer, and a history of polyposis  
  - Has a PTEN mutation and NOT uterine cancer  
  - Has a TP53 mutation AND either personal history of breast cancer diagnosed <65 years old, OR personal history of any other cancer diagnosed <40 years old  
  - Has a RB1 mutation with personal/family history of retinoblastoma or other associated RB tumor  
  - Has a RET mutation with personal history of medullary thyroid cancer/family history of thyroid cancer

*require additional vetting per protocol

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**Treatment Plan**

See Section 5.0 for Complete Methodology Details

**Primary Intervention Study (RCT) (Step 1):**

- Genetic education website (Arm A—intervention): Access to the web-based genetic education module at the study site following completion of the baseline survey; will be able to revisit the site using any computer with web access, including a smart phone
- Usual care (Arm B—control group): Conversations with treating physicians, interaction with/information from the MATCH research staff, additional information that the patient seeks out through usual resources; 3 months after receipt of results, these patients will be provided access to the web-based education module
- Patients on both arms will be asked to complete 4 surveys electronically

**Secondary Genetic Counseling Substudy (Step 2—Arms D & E; potential germline mutation patients):**

- Genetic counseling: Remote visits and surveys (i.e., visit 1- pre-test counseling, visit 2- result disclosure/review of implications for relatives)
- Germline genetic testing: Genetic counselor to provide recommendations for testing; treating physician to order the germline testing
- Will complete up to 4 surveys/associative interventions

**Alternative Pathway to Substudy (Step 1—Arm C, then Step 2—Arm E):**

- For a subgroup of MATCH patients who receive their tumor molecular profile results and meet the criteria for potential actionable germline mutation; not required to complete the main study (Step 1)

**See protocol for the PRO/QOL Data Collection Process and Study Instruments**

Duration of participation: ~4-5 months for Primary Intervention Study; ~1-4 months for Substudy (Step 2), depending on if patients have confirmatory germline testing or not; ≤ 5 months if participating on both the study and substudy

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**Patient Enrollment**

All Sites: Oncology Patient Enrollment Network (OPEN) [https://open.ctsu.org](https://open.ctsu.org)

**Protocol Information**

ECOG-ACRIN Operations-Boston: 857-504-2900, [http://ecog-acrin.org](http://ecog-acrin.org) (Member Login)

Please Enroll Your Eligible Patients!
Schema

Step 1 - Primary Online Genetic Education Study (RCT)

1. Patient consented to COMET RCT, patient enrolled in MATCH screening (Step 0), and proceeded with biopsy and molecular profiling.
2. Randomized to Arm A or B
3. Post-education survey (T0): 1-2 weeks after completion of T0
4. Screening participant (patient) receives treatment assignment results and/or tumor molecular profile results from their research/clinical team/treating physician.
5. Post-disclosure survey (T2): Within 3 days from receipt of results.
6. Usual care (Arm B) participants provided access to online genetic education.
7. Additional participation in the secondary genetic counseling substudy.

Step 2 - Secondary Genetic Counseling Substudy

1. Patient participating in COMET RCT (Step 1: online genetic education) is interested in pursuing Step 2 after receiving their tumor testing results.
2. Patient meets criteria for potential actionable genetic mutation based on their testing results, and other eligibility requirements.
3. Patient consents to GC Substudy.
4. Baseline survey (T0SS):
   - For Arm D participants: Limited items (T0SS populated with T2 information)
   - For Arm E participants: Baseline survey
5. If patient agrees to confirmatory testing:
   - GC Visit 2: Post V2 survey (T2SS): Within 3 days of GC Visit 2
   - 3 month survey (T3SS): 3 months after GC Visit 2

Accrual Goal (Primary Online Genetic Education Study): 670

1. Select expansion sites only; see Section 4.1, 4.5 for more information.
2. Stratification Factors:
   a. Gender (Male vs. Female)
   b. Race (White vs. Others)
   c. Age (<65 vs. >65 years)
   d. Education level: No College vs. College vs. Professional/Graduate
3. If patient is eligible for COMET Step 2 (secondary genetic counseling substudy) based on their testing results and the eligibility criteria, they can also continue to participate in Step 2 in parallel with Step 1. The 3 month survey (T3) for Step 1 will be still be requested, even if Step 2 is pursued.

Accrual Goal (Secondary Genetic Counseling Study): 100

4. Select Participating Institutions Only; see Section 4.2.2 for instructions and requirements.
5. Please note, per Section 3.2.1, 1.1, patients entering the study via an expansion site (see footnote #1 in the Step 1 schema) are only eligible for Step 2 if they participated in Step 1. They are not eligible for Step 2 via this path, without any prior participation in the study.
6. GC = genetic counseling
7. SS = substudy