

# Oncology Clinical Pathways

## Colon Cancer

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April 2024 – V3.2024



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U.S. Department  
of Veterans Affairs

# Table of Contents

<a href="#">Presumptive Conditions</a> .....	3
<a href="#">Stage I</a> .....	4
<a href="#">Stage II</a> .....	5
<a href="#">Stage III</a> .....	6
<a href="#">Stage IV Initial Evaluation</a> .....	7
<a href="#">Stage IV Potentially Resectable</a> .....	8
<a href="#">Stage IV Unresectable, pMMR First Line, Fit for Chemotherapy</a> .....	9
<a href="#">Stage IV Unresectable, pMMR Second Line, Fit for Chemotherapy</a> .....	10
<a href="#">Stage IV Unresectable, pMMR Third Line, Fit for Chemotherapy</a> .....	11
<a href="#">Stage IV Unresectable, pMMR Additional Lines, Fit for Chemotherapy</a> .....	12
<a href="#">Stage IV Unresectable, dMMR</a> .....	13
<a href="#">Stage IV Unresectable, Unfit for Combination Chemotherapy</a> .....	14
<a href="#">Molecular Testing</a> .....	15
<a href="#">Molecular Testing Table</a> .....	16



# Colon Cancer – Presumptive Conditions

VA automatically presumes that certain disabilities were caused by military service. This is because of the unique circumstances of a specific Veteran's military service. If a presumed condition is diagnosed in a Veteran within a certain group, they can be awarded disability compensation.

## Atomic Veterans – Exposure to Ionizing Radiation

- Cancer of the colon

## Gulf War and Post 9/11 Veterans

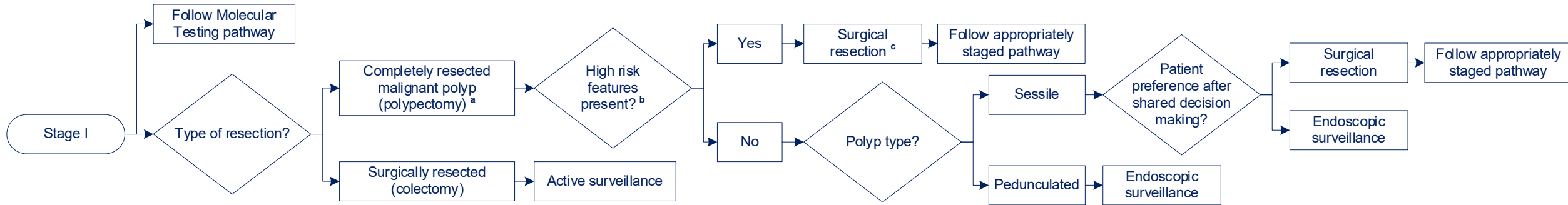
If the patient served on or after Sept. 11, 2001, in Afghanistan, Djibouti, Egypt, Jordan, Lebanon, Syria, Uzbekistan, or Yemen or if you served in the \*Southwest Asia theater of operations, or Somalia, on or after Aug. 2, 1990, specific conditions include:

- Gastrointestinal cancer of any type

\* The Southwest Asia theater of operations refers to Iraq, Kuwait, Saudi Arabia, the neutral zone between Iraq and Saudi Arabia, Bahrain, Qatar, the United Arab Emirates, Oman, the Gulf of Aden, the Gulf of Oman, the Persian Gulf, the Arabian Sea, the Red Sea, and the airspace above these locations.

For more information, please visit [U.S. Department of Veterans Affairs - Presumptive Disability Benefits \(va.gov\)](https://www.va.gov)

# Colon Cancer – Stage I



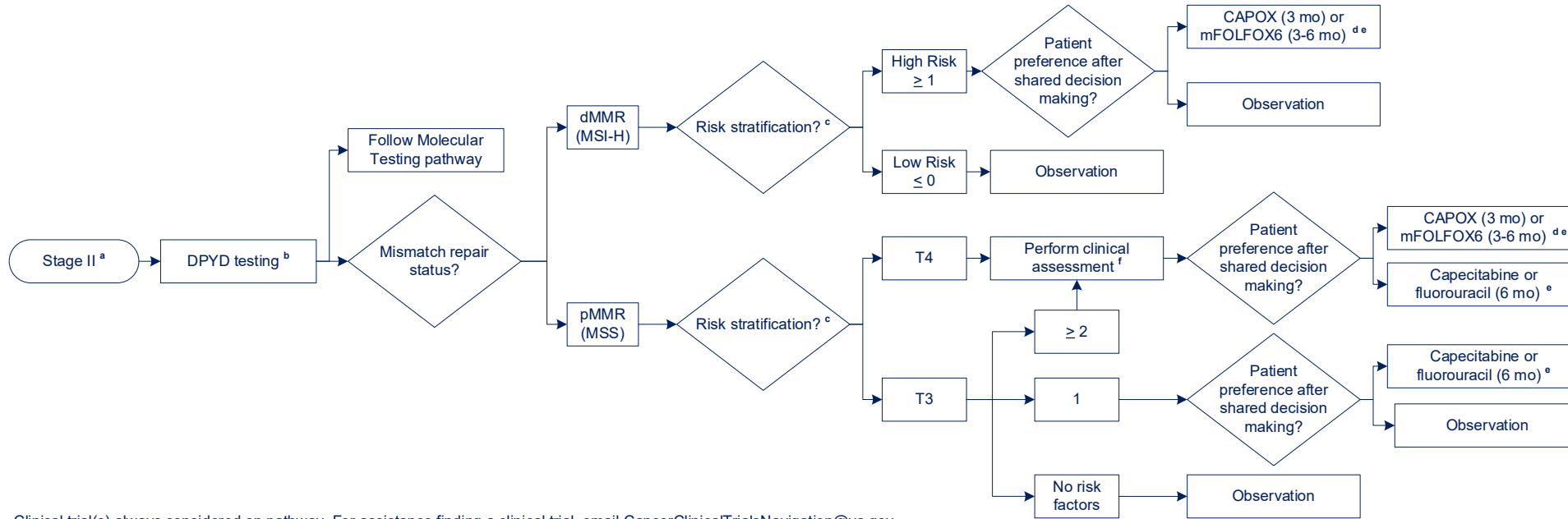
Clinical trial(s) always considered on pathway. For assistance finding a clinical trial, email [CancerClinicalTrialsNavigation@va.gov](mailto:CancerClinicalTrialsNavigation@va.gov).

<sup>a</sup> **Resection** if not a complete resection, additional endoscopic resection may be considered if feasible

<sup>b</sup> **High Risk Features for Sessile Polyps** includes: poor tumor differentiation, lymphovascular invasion, submucosal invasion depth >1 mm, tumor involvement of the cautery margin, tumor budding; **High Risk Features for Pedunculated Polyps** includes: poor tumor differentiation, lymphovascular invasion, tumor within 1 mm of the resection margin

<sup>c</sup> **Surgical Resection** defined as a formal oncologic segmental colectomy

# Colon Cancer – Stage II



Clinical trial(s) always considered on pathway. For assistance finding a clinical trial, email [CancerClinicalTrialsNavigation@va.gov](mailto:CancerClinicalTrialsNavigation@va.gov).

<sup>a</sup> **Stage II** determined following surgical resection, with time to initiation of chemotherapy recommended to begin 6-8 weeks following surgery; delay beyond 3 months is likely to greatly reduce the efficacy of subsequent chemotherapy and should be weighed into decision making; late initiation of chemotherapy should be evaluated on a case by case basis

<sup>b</sup> **Perform DPYD Testing If Not Already Performed** if DPYD PGx results return predicted phenotypes of either intermediate or poor metabolizer, please consult your local PGx pharmacist or submit an IFC Pharmacogenomics e-consult for assistance with therapeutic recommendation; a clinician may proceed without DPYD results if withholding chemotherapy for 2-3 weeks may gravely endanger patient's life; for example, if the disease burden is very high and it involves a large portion of vital organs such as liver, etc.

<sup>c</sup> **Risk Stratification:** high risk features include presence of clinical obstruction, localized tumor perforation, or poorly differentiated tumor (pMMR only), lymphovascular invasion, perineural invasion, <12 harvested lymph nodes, positive and/or close margin ( $\leq 1$ mm in non-peritonealized margins), or high tumor budding (18/0.785 mm<sup>2</sup>)

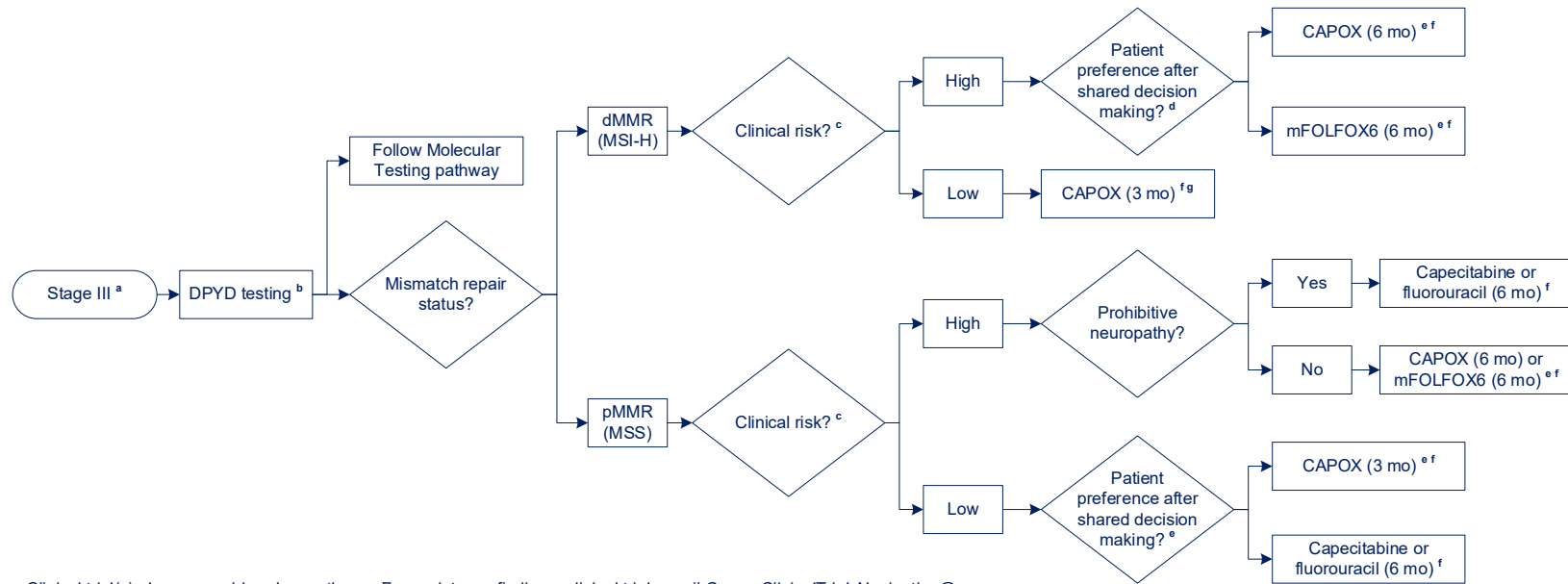
<sup>d</sup> **Oxaliplatin-Based Regimens** risk of  $\geq$  Grade 3 neurotoxicity is lower with 3-month vs. 6-month; if  $\geq$  grade 2 neuropathy develops during treatment, may discontinue oxaliplatin after three months while continuing fluoropyrimidine to full course (6 months); for pMMR, the benefit of addition of oxaliplatin is unclear in patients aged > 70 due to paucity of data

<sup>e</sup> **Capecitabine** avoid capecitabine if adherence issues, unable to self-report toxicity, or severe renal impairment (CrCl < 30ml/min)

<sup>f</sup> **Clinical Assessment** determine presence of renal dysfunction, comorbidities, neuropathy, limited performance status, or limited life expectancy, as they would negatively impact the decision to use adjuvant chemotherapy

DPYD Dihydropyrimidine Dehydrogenase  
dMMR Mismatch Repair Deficient  
MMR Mismatch Repair  
MSI Microsatellite Instability  
MSS Microsatellite Stable  
pMMR Proficient Mismatch Repair

# Colon Cancer – Stage III



Clinical trial(s) always considered on pathway. For assistance finding a clinical trial, email [CancerClinicalTrialsNavigation@va.gov](mailto:CancerClinicalTrialsNavigation@va.gov).

<sup>a</sup> **Stage III** determined following surgical resection, with time to initiation of chemotherapy recommended to begin 6-8 weeks following surgery; delay beyond 3 months is likely to greatly reduce the efficacy of subsequent chemotherapy and should be weighed into decision making; late initiation of chemotherapy should be evaluated on a case by case basis

<sup>b</sup> **Perform DPYD Testing If Not Already Performed** if DPYD PGx results return predicted phenotypes of either intermediate or poor metabolizer, please consult your local PGx pharmacist or submit an IFC Pharmacogenomics e-consult for assistance with therapeutic recommendation; a clinician may proceed without DPYD results if withholding chemotherapy for 2-3 weeks may gravely endanger patient's life; for example, if the disease burden is very high and it involves a large portion of vital organs such as liver, etc.

<sup>c</sup> **Clinical Risk** defines high risk as T4 and/or N2 ( $\geq 4$  positive nodes) and low risk as T1-3 and N1

<sup>d</sup> **If Preexisting Neuropathy**, single agent treatment may be an alternative for pMMR tumor; however, fluorouracil/capecitabine single agent treatment represents an inadequate option for dMMR tumors; observation may be appropriate for dMMR with significant baseline neuropathy ( $\geq$  grade 2)

<sup>e</sup> **Oxaliplatin-Based Regimens** risk of  $\geq$  grade 3 neurotoxicity is lower with 3-month vs. 6-month; if  $>$  grade 2 neuropathy develops, discontinue oxaliplatin while continuing fluoropyrimidine to full course; benefit of adding oxaliplatin is unclear in patients aged  $>$  70 due to paucity of data

<sup>f</sup> **Capecitabine** avoid capecitabine if adherence issues, unable to self-report toxicity, or severe renal impairment (CrCl  $<$  30ml/min)

**DPYD** Dihydropyrimidine Dehydrogenase

**dMMR** Mismatch Repair Deficient

**MMR** Mismatch Repair

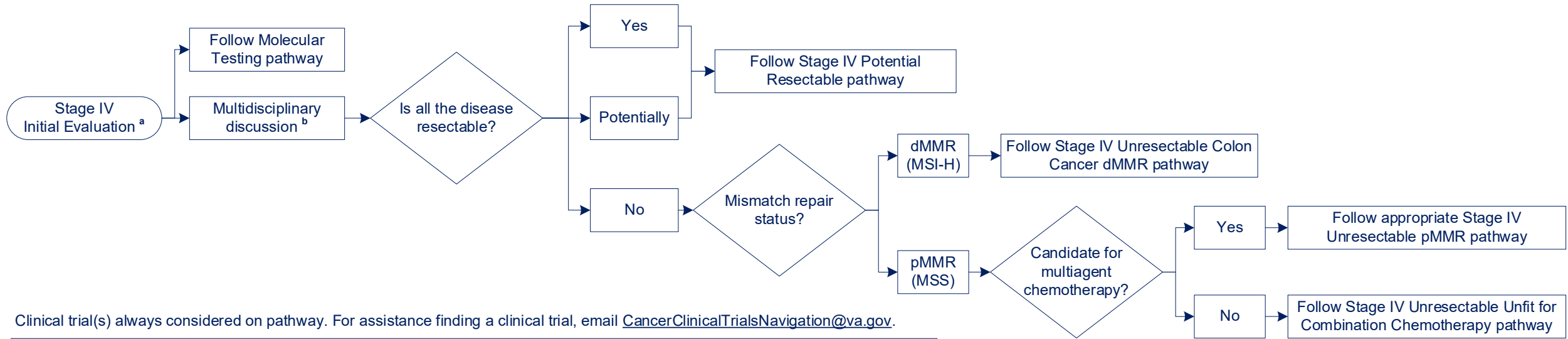
**MSI** Microsatellite Instability

**MSI-H** Microsatellite Instability High

**MSS** Microsatellite Stable

**pMMR** Proficient Mismatch Repair

# Colon Cancer – Stage IV Initial Evaluation



Clinical trial(s) always considered on pathway. For assistance finding a clinical trial, email [CancerClinicalTrialsNavigation@va.gov](mailto:CancerClinicalTrialsNavigation@va.gov).

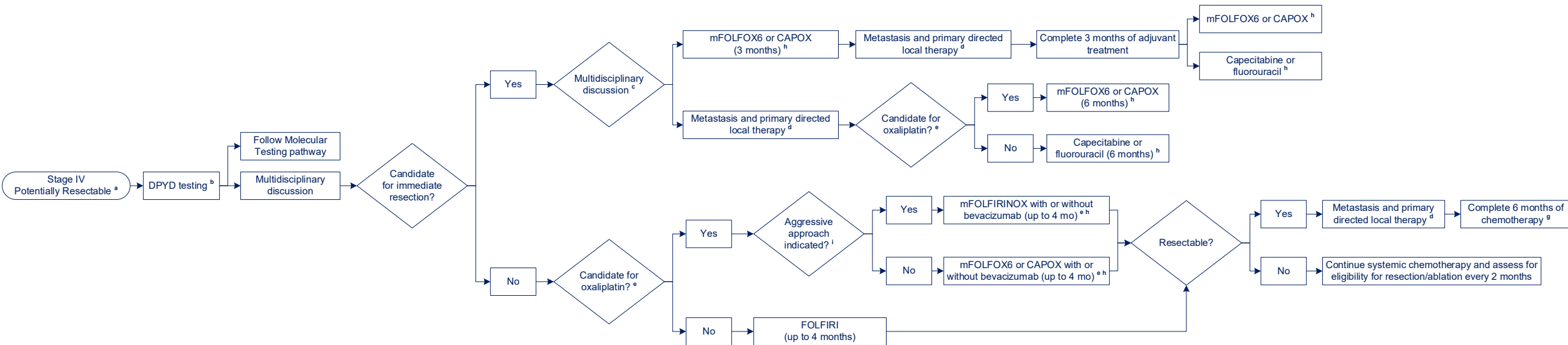
<sup>a</sup> **Stage IV** determined by clinical evaluation and imaging and/or pathological confirmation post diagnosis

<sup>b</sup> **Multidisciplinary Discussion** refers to tumor board or with an expert or group of experts including but not limited to Medical Oncology, Radiation Oncology, Surgical Oncology, Thoracic Surgery, Interventional Radiology, Diagnostic Radiology, and/or Pathology

**dMMR** Mismatch Repair Deficient  
**MMR** Mismatch Repair  
**MSI-H** Microsatellite Instability High  
**MSS** Microsatellite Stable  
**pMMR** Proficient Mismatch Repair



# Colon Cancer – Stage IV Potentially Resectable



Clinical trial(s) always considered on pathway. For assistance finding a clinical trial, email [CancerClinicalTrialsNavigation@va.gov](mailto:CancerClinicalTrialsNavigation@va.gov).

<sup>a</sup> If Neuropathy  $\geq$  Grade 2 Develops during neoadjuvant phase, complete treatment with fluorouracil and capecitabine

<sup>b</sup> Perform DPYD Testing if Not Already Performed if DPYD PGx results return predicted phenotypes of either intermediate or poor metabolizer, please consult your local PGx pharmacist or submit an IFC Pharmacogenomics e-consult for assistance with therapeutic recommendation; a clinician may proceed without DPYD results if withholding chemotherapy for 2-3 weeks may gravely endanger patient's life; for example, if the disease burden is very high and it involves a large portion of vital organs such as liver, etc.

<sup>c</sup> Multidisciplinary Discussion refers to tumor board or with an expert or group of experts including but not limited to Medical Oncology, Radiation Oncology, Surgical Oncology, Thoracic Surgery, Interventional Radiology, Diagnostic Radiology, and/or Pathology

<sup>d</sup> Metastasis-Directed Local Therapy options include surgery, radiation, and IR ablative techniques; surgery is preferred if feasible; resection of primary if present

<sup>e</sup> Candidate for Oxaliplatin contraindication if any adjuvant treatment in the past 12 months or preexisting neuropathy  $>1$  grade neuropathy

<sup>f</sup> Candidate for Bevacizumab received fluoropyrimidine and platinum agent in the first-line setting; ECOG PS 0-2; ANC  $\geq$  1500/mm<sup>3</sup>; due to anti-VEGF effects patients with the following should not receive bevacizumab: non-healing wound/fracture, major surgery in prior 4 weeks, bleeding disorder or coagulopathy, recent history of GI perforation, unstable cardiac condition (uncontrolled HTN, arterial thromboembolism, symptomatic CHF (NYHA II-IV) or arrhythmia), or active cocaine use

<sup>g</sup> Choice of Chemotherapy will be between oxaliplatin-based doublet (if eligible for oxaliplatin) or single agent capecitabine and fluorouracil; if neuropathy  $\geq$  grade 2 develop during neoadjuvant phase, complete treatment with capecitabine and fluorouracil

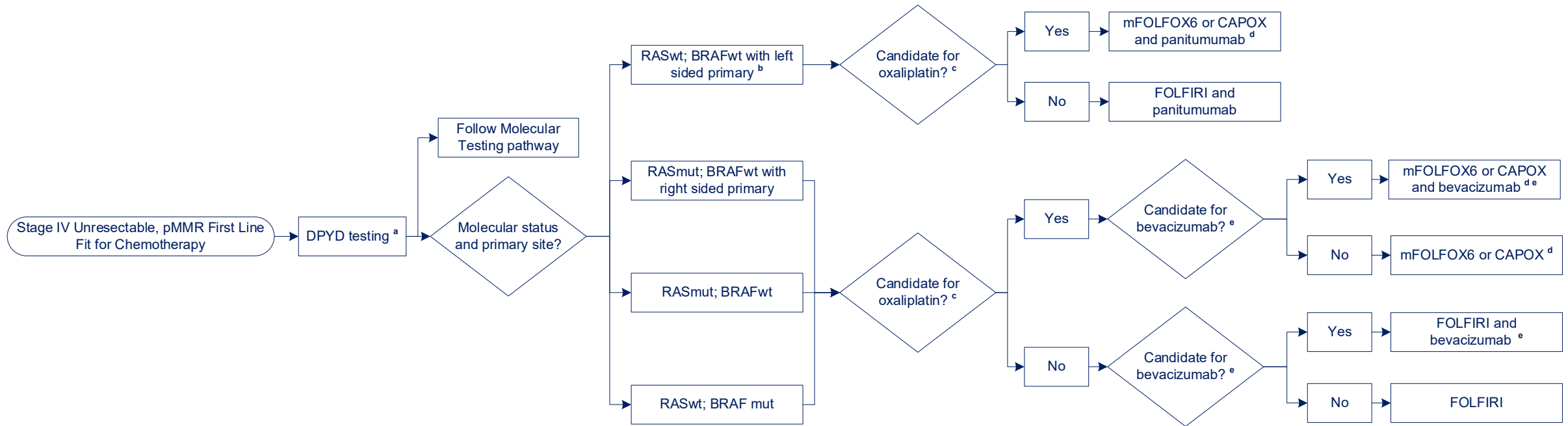
<sup>h</sup> Capecitabine avoid capecitabine if adherence issues, unable to self-report toxicity, or severe renal impairment (CrCl  $<$  30ml/min)

<sup>i</sup> Aggressive Approach Indicated may be considered in very fit patients with excellent performance status and high disease burden and/or presence of BRAF mutation

DPYD Dihydropyrimidine Dehydrogenase



# Colon Cancer – Stage IV Unresectable, pMMR First Line Fit for Chemotherapy



Clinical trial(s) always considered on pathway. For assistance finding a clinical trial, email [CancerClinicalTrialsNavigation@va.gov](mailto:CancerClinicalTrialsNavigation@va.gov).

<sup>a</sup> **Perform DPYD Testing If Not Already Performed** if DPYD PGx results return predicted phenotypes of either intermediate or poor metabolizer, please consult your local PGx pharmacist or submit an IFC Pharmacogenomics e-consult for assistance with therapeutic recommendation; a clinician may proceed without DPYD results if withholding chemotherapy for 2-3 weeks may gravely endanger patient's life; for example, if the disease burden is very high and it involves a large portion of vital organs such as liver, etc.

<sup>b</sup> **Left Sided Primary** is defined as primary originating in splenic flexure and colon distal to that

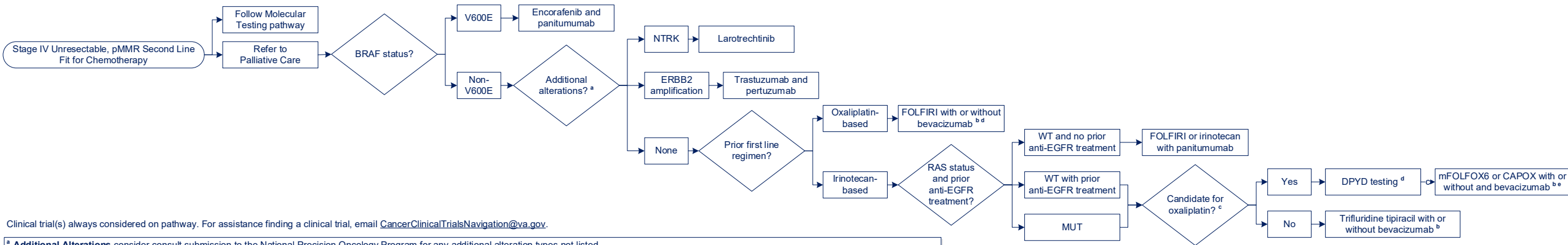
<sup>c</sup> **Candidate for Oxaliplatin** contraindication if any adjuvant treatment in the past 12 months or preexisting neuropathy >1 grade neuropathy; patient preference to avoid neuropathy

<sup>d</sup> **Capecitabine** avoid capecitabine if adherence issues, unable to self-report toxicity, or severe renal impairment (CrCl <30 ml/min)

<sup>e</sup> **Candidate for Bevacizumab** received fluoropyrimidine and platinum agent in the first-line setting; ECOG PS 0-2; ANC  $\geq$  1500/mm<sup>3</sup>; due to anti-VEGF effects patients with the following should not receive bevacizumab: non-healing wound/fracture, major surgery in prior 4 weeks, bleeding disorder or coagulopathy, recent history of GI perforation, unstable cardiac condition (uncontrolled HTN, arterial thromboembolism, symptomatic CHF (NYHA II-IV) or arrhythmia), or active cocaine use

DPYD Dihydropyrimidine Dehydrogenase  
 mut mutation  
 pMMR Proficient Mismatch Repair  
 wt wild type

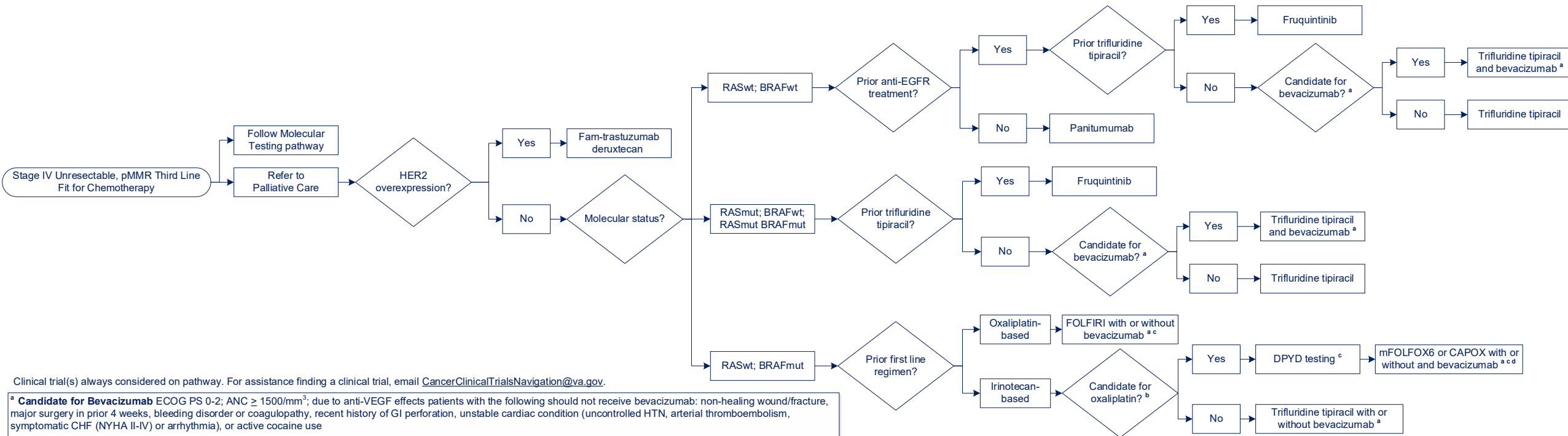
# Colon Cancer – Stage IV Unresectable, pMMR Second Line Fit for Chemotherapy



Clinical trial(s) always considered on pathway. For assistance finding a clinical trial, email [CancerClinicalTrialsNavigation@va.gov](mailto:CancerClinicalTrialsNavigation@va.gov).

- <sup>a</sup> **Additional Alterations** consider consult submission to the National Precision Oncology Program for any additional alteration types not listed
  - <sup>b</sup> **Candidate for Bevacizumab** received fluoropyrimidine and platinum agent in the first-line setting; ECOG PS 0-2; ANC  $\geq 1500/\text{mm}^3$ ; due to anti-VEGF effects patients with the following should **not** receive bevacizumab: non-healing wound/fracture, major surgery in prior 4 weeks, bleeding disorder or coagulopathy, recent history of GI perforation, unstable cardiac condition (uncontrolled HTN, arterial thromboembolism, symptomatic CHF (NYHA II-IV) or arrhythmia), or active cocaine use
  - <sup>c</sup> **Candidate for Oxaliplatin** contraindication if disease progression within 12 months of adjuvant treatment or preexisting neuropathy >1 grade neuropathy
  - <sup>d</sup> **Perform DPYD Testing If Not Already Performed** if DPYD PGx results return predicted phenotypes of either intermediate or poor metabolizer, please consult your local PGx pharmacist or submit an IFC Pharmacogenomics e-consult for assistance with therapeutic recommendation; a clinician may proceed without DPYD results if withholding chemotherapy for 2-3 weeks may gravely endanger patient's life; for example, if the disease burden is very high and it involves a large portion of vital organs such as liver, etc.
  - <sup>e</sup> **Capecitabine** avoid capecitabine if adherence issues, unable to self-report toxicity, or severe renal impairment (CrCl <30 ml/min)
- DPYD** Dihydropyrimidine Dehydrogenase  
**EGFR** Epidermal Growth Factor Receptor  
**MUT** Mutation  
**NTRK** Neurotrophic Tyrosine Receptor Kinase  
**pMMR** Proficient Mismatch Repair  
**WT** Wild Type

# Colon Cancer – Stage IV Unresectable, pMMR Third Line Fit for Chemotherapy



Clinical trial(s) always considered on pathway. For assistance finding a clinical trial, email [CancerClinicalTrialsNavigation@va.gov](mailto:CancerClinicalTrialsNavigation@va.gov).

<sup>a</sup> **Candidate for Bevacizumab** ECOG PS 0-2; ANC  $\geq$  1500/mm<sup>3</sup>; due to anti-VEGF effects patients with the following should not receive bevacizumab: non-healing wound/fracture, major surgery in prior 4 weeks, bleeding disorder or coagulopathy, recent history of GI perforation, unstable cardiac condition (uncontrolled HTN, arterial thromboembolism, symptomatic CHF (NYHA II-IV) or arrhythmia), or active cocaine use

<sup>b</sup> **Capecitabine or Fluorouracil** consider DPYD testing to inform starting dose of fluorouracil or capecitabine in the shared decision making

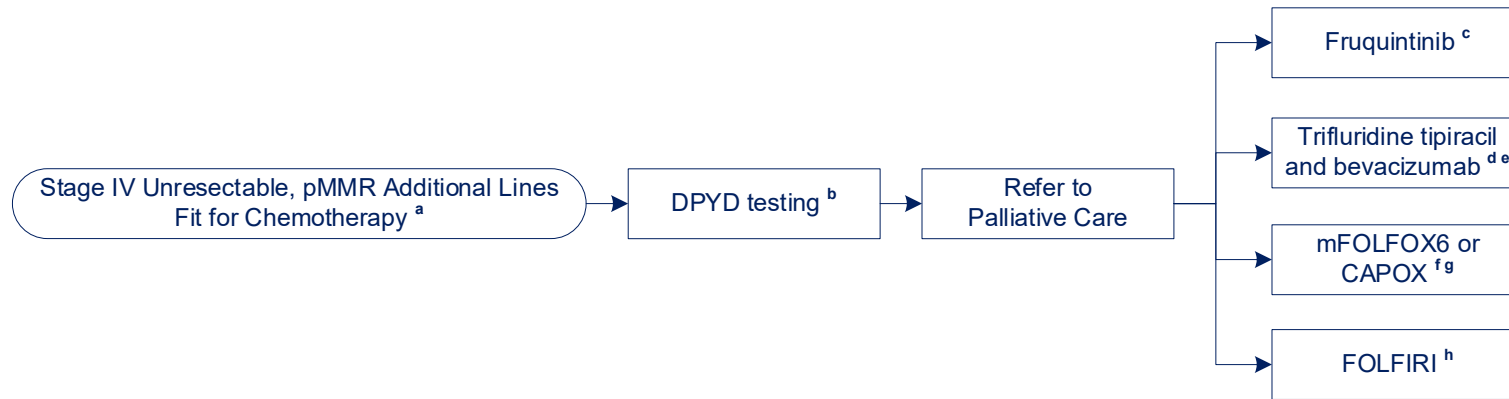
<sup>c</sup> **Perform DPYD Testing If Not Already Performed** if DPYD PGx results return predicted phenotypes of either intermediate or poor metabolizer, please consult your local PGx pharmacist or submit an IFC Pharmacogenomics e-consult for assistance with therapeutic recommendation; a clinician may proceed without DPYD results if withholding chemotherapy for 2-3 weeks may gravely endanger patient's life; for example, if the disease burden is very high and it involves a large portion of vital organs such as liver, etc.

<sup>d</sup> **Capecitabine** avoid capecitabine if adherence issues, unable to self-report toxicity, or severe renal impairment (CrCl <30 ml/min)

DPYD Dihydropyrimidine Dehydrogenase  
 EGFR Epidermal Growth Factor Receptor  
 mut mutation  
 pMMR Proficient Mismatch Repair  
 wt wild type



# Colon Cancer – Stage IV Unresectable, pMMR Additional Lines Fit for Chemotherapy



Clinical trial(s) always considered on pathway. For assistance finding a clinical trial, email [CancerClinicalTrialsNavigation@va.gov](mailto:CancerClinicalTrialsNavigation@va.gov).

<sup>a</sup> **Performance Status** for a patient with performance status from 0-2

<sup>b</sup> **Perform DPYD Testing If Not Already Performed** if DPYD PGx results return predicted phenotypes of either intermediate or poor metabolizer, please consult your local PGx pharmacist or submit an IFC Pharmacogenomics e-consult for assistance with therapeutic recommendation; a clinician may proceed without DPYD results if withholding chemotherapy for 2-3 weeks may gravely endanger patient's life; for example, if the disease burden is very high and it involves a large portion of vital organs such as liver, etc.

<sup>c</sup> **Fruquintinib** no prior failure to the treatment

<sup>d</sup> **Trifluridine Tipiracil** no prior failure to the treatment

<sup>e</sup> **Candidate for Bevacizumab** received fluoropyrimidine and platinum agent in the first-line setting; ECOG PS 0-2; ANC  $\geq 1500/\text{mm}^3$ ; due to anti-VEGF effects patients with the following should not receive bevacizumab: non-healing wound/fracture, major surgery in prior 4 weeks, bleeding disorder or coagulopathy, recent history of GI perforation, unstable cardiac condition (uncontrolled HTN, arterial thromboembolism, symptomatic CHF (NYHA II-IV) or arrhythmia), or active cocaine use

<sup>f</sup> **FOLFOX6 or CAPOX** well tolerated if used previously and duration from last treatment >12 months and and no  $\geq$  grade 2 neuropathy

<sup>g</sup> **Capecitabine** avoid capecitabine if adherence issues, unable to self-report toxicity, or severe renal impairment (CrCl <30 ml/min)

<sup>h</sup> **FOLFIRI** well tolerated if used previously and duration from last treatment >12 months

**DPYD** Dihydropyrimidine Dehydrogenase

**pMMR** Proficient Mismatch Repair

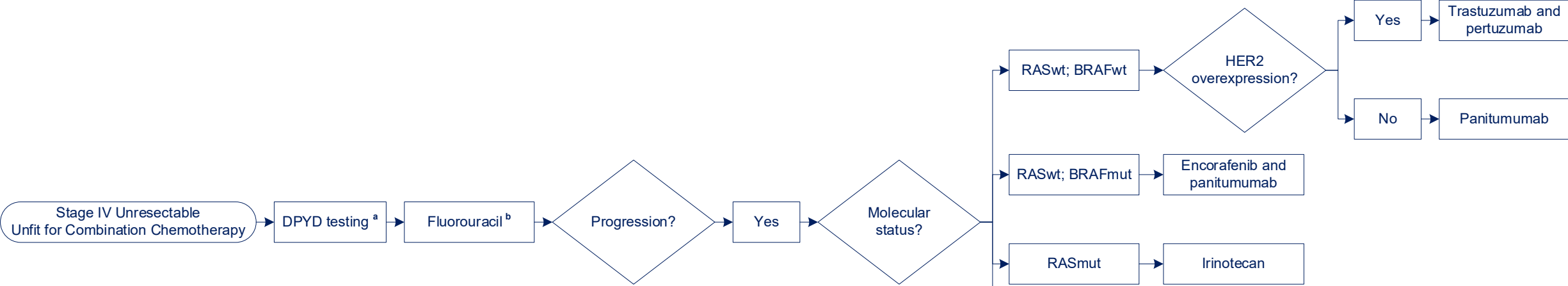
# Colon Cancer – Stage IV Unresectable dMMR



Clinical trial(s) always considered on pathway. For assistance finding a clinical trial, email [CancerClinicalTrialsNavigation@va.gov](mailto:CancerClinicalTrialsNavigation@va.gov).

**dMMR** Mismatch Repair Deficient  
**pMMR** Proficient Mismatch Repair

# Colon Cancer – Stage IV Unresectable Unfit for Combination Chemotherapy



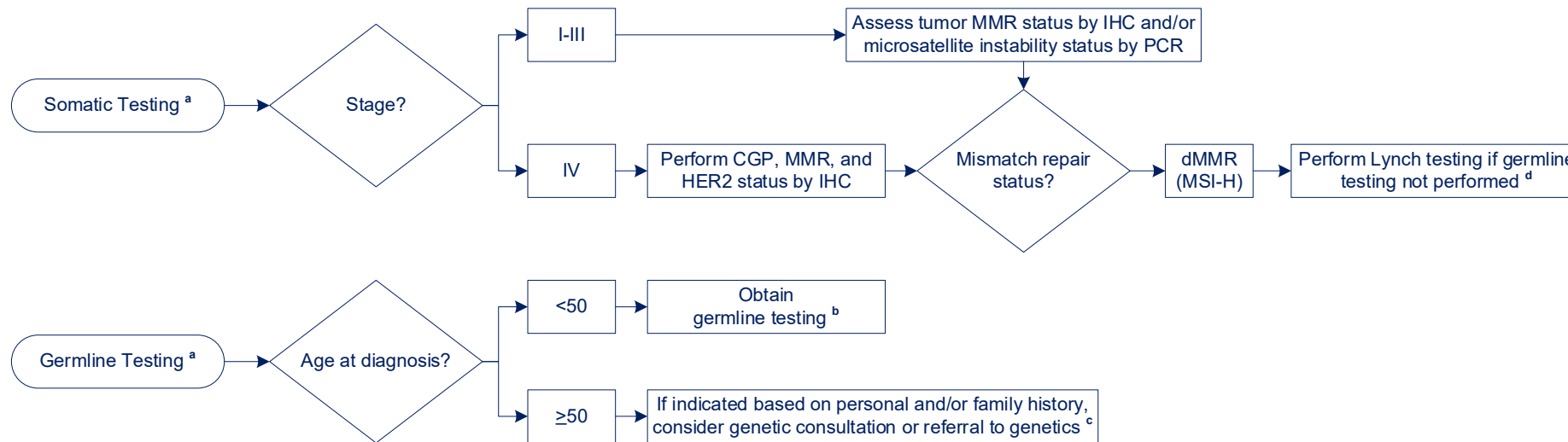
Clinical trial(s) always considered on pathway. For assistance finding a clinical trial, email [CancerClinicalTrialsNavigation@va.gov](mailto:CancerClinicalTrialsNavigation@va.gov).

<sup>a</sup> **Perform DPYD Testing If Not Already Performed** if DPYD PGx results return predicted phenotypes of either intermediate or poor metabolizer, please consult your local PGx pharmacist or submit an IFC Pharmacogenomics e-consult for assistance with therapeutic recommendation; a clinician may proceed without DPYD results if withholding chemotherapy for 2-3 weeks may gravely endanger patient's life; for example, if the disease burden is very high and it involves a large portion of vital organs such as liver, etc.

<sup>b</sup> **Fluorouracil** infusional 5-fluorouracil monotherapy is preferred, however for select patient unable to tolerate pump, capecitabine may be substituted if no issues with adherence, toxicity reporting, or severe renal impairment (CrCl<30 ml.min)

**DPYD** Dihydropyrimidine Dehydrogenase  
**mut** mutation  
**wt** wild type

# Colon Cancer – Molecular Testing



Clinical trial(s) always considered on pathway. For assistance finding a clinical trial, email [CancerClinicalTrialsNavigation@va.gov](mailto:CancerClinicalTrialsNavigation@va.gov).

<sup>a</sup> **Molecular Testing** perform for pathologically confirmed cancer

<sup>b</sup> **Germline Testing** an appropriate germline testing panel should include at minimum the following genes: APC; AXIN2; BMPR1A; CHEK2; EPCAM; GALNT12; GREM1; MLH1; MLH3; MSH2; MSH3; MSH6; MUTYH; NTHL1; PMS2; POLD1; POLE; PTEN; RNF43; SMAD4; STK11; and TP53

<sup>c</sup> **Personal and Family History** consider germline testing if significant personal and/or family history of multiple polyps, other Lynch syndrome or other hereditary cancer syndrome-associated cancers (e.g., colorectal, endometrial, gastric, ovarian, pancreas, urothelial, brain (usually glioblastoma), biliary tract, and small intestine, as well as sebaceous adenomas, sebaceous carcinomas, and keratoacanthomas as seen in Muir-Torre syndrome), or pathogenic or likely pathogenic variant in a gene associated with known hereditary cancer syndrome is present in the patient or a family member

<sup>d</sup> **Lynch Testing** the diagnostic Lynch genetic testing algorithm depends on the pattern of MLH1, MSH2, MSH6, and PMS2 expression by IHC; diagnostic Lynch genetic testing should be performed if there is loss of MSH2, MSH6, MSH2/MSH6, or PMS2 expression by IHC; if there is loss of MLH1 expression by IHC, *MLH1* promoter hypermethylation testing should be performed; *MLH1* hypermethylation suggests somatic mutation, but diagnostic Lynch genetic testing should be performed if *MLH1* is unmethylated in the context of MLH1 loss by IHC; a diagnostic Lynch genetic testing panel should include at minimum the following genes: *EPCAM*, *MLH1*, *MSH2*, *MSH6*, and *PMS2*

**CGP** Comprehensive Genomic Profiling

**dMMR** Mismatch Repair Deficient

**IHC** Immunohistochemistry

**MMR** Mismatch Repair

**MSI-H** Microsatellite Instability High

**PCR** Polymerase Chain Reaction



# Colon Cancer – Molecular Testing Table

Eligibility	Test Category	Test Type	Recommended Vendors	NPOP Coverage	Specimen Type
Stage I-III	IHC*	Mismatch repair (MMR) protein expression by IHC: MLH1, MSH2, MSH6, and PMS2.	Local VA or locally contracted vendor	No	Tumor Tissue
	PCR*	Microsatellite instability (MSI) status by PCR.	Regional VA Testing Center (GLA)	Yes	Tumor Tissue and Normal Tissue or Blood
	IHC	Consider BRAF V600E IHC if MLH1 or PMS2 expression is lost by IHC, or if MSI-H and IHC not performed. Mutated suggests somatic mutation. Unmutated calls for Methylation testing.	Local VA or locally contracted vendor	No	Tumor Tissue
	Molecular Testing	Consider BRAF V600E mutation testing if MLH1 or PMS2 expression is lost by IHC, or if MSI-H and IHC not performed. Mutated suggests somatic mutation. Unmutated calls for Methylation testing.	Local VA or locally contracted vendor	No	Tumor Tissue
	Methylation Testing	MLH1 promoter hypermethylation testing (in the setting of loss of MLH1 or PMS2 expression by IHC). Hypermethylation suggests somatic mutation. Unmethylated calls for Germline Lynch testing.	Local VA or locally contracted vendor	No	Tumor Tissue
	Germline NGS***	If full germline testing not performed, perform Germline Lynch testing if: 1) MSH2 or MSH6 loss by IHC or 2) MLH1 or PMS2 loss by IHC and MLH1 unmethylated or 3) MSI-H without IHC testing AND BRAF unmutated AND MLH1 unmethylated	Fulgent Prevention Genetics	Yes Yes	Saliva, Blood
Stage IV	Somatic NGS	Comprehensive genomic profiling (CGP) including MSI.	Tempus Foundation Medicine	Yes Yes	Tumor Tissue****, Blood
	IHC	HER2	Local VA or locally contracted vendor	No	Tumor Tissue
	FISH	Reflex to HER2 FISH if 2+ on IHC	Local VA or locally contracted vendor	No	Tumor Tissue
	IHC**	Mismatch repair (MMR) protein expression by IHC: MLH1, MSH2, MSH6, and PMS2.	Tempus	Yes (When ordered with CGP)	Tumor Tissue
	PCR**	Consider microsatellite instability (MSI) status by PCR if MSI by CGP is not performed or equivocal.	Regional VA Testing Center (GLA)	Yes	Tumor Tissue and Normal Tissue or Blood
	Methylation Testing	MLH1 promoter hypermethylation testing (in the setting of loss of MLH1 or PMS2 expression by IHC). Hypermethylation suggests somatic mutation. Unmethylated calls for Lynch testing.	Local VA or locally contracted vendor	No	Tumor Tissue
	Germline NGS***	If full germline testing not performed, perform Germline Lynch testing if: 1) MSH2 or MSH6 loss by IHC or 2) MLH1 or PMS2 loss by IHC and MLH1 unmethylated or 3) MSI-H without IHC testing AND BRAF unmutated AND MLH1 unmethylated	Fulgent Prevention Genetics	Yes Yes	Saliva, Blood
Diagnosis Below the Age of 50	Germline NGS****	Full Germline Testing	Fulgent Prevention Genetics	Yes Yes	Saliva, Blood
Personal and/or Family History of Multiple Polyps, Other Lynch Syndrome or Other Hereditary Cancer Syndrome Associated Cancers, or Pathogenic or Likely Pathogenic Variant in a Gene Associated with Known Hereditary Cancer Syndrome is Present in the Patient or a Family Member	Germline NGS****	Full Germline Testing	Fulgent Prevention Genetics	Yes Yes	Saliva, Blood

\* For Stage I-III, either MMR or MSI or Both can be performed

\*\* For Stage IV, Both MMR and MSI should be performed; If MSI cannot be determined by CGP, then MSI by PCR can be performed

\*\*\* Germline Lynch testing should include at minimum the following genes: EPCAM (deletion), MLH1, MSH2, MSH6, PMS2, POLE, and POLD1

\*\*\*\* VA Common Hereditary POC panel or Equivalent Germline Test; Full Germline testing should include at minimum the following genes: APC; AXIN2; BMPR1A; CHEK2; EPCAM; GALNT12; GREM1; MLH1; MLH3; MSH2; MSH3; MSH6; MUTYH; NTHL1; PMS2; POLD1; POLE; PTEN; RNF43; SMAD4; STK11; and TP53; For genetic online ordering, refer to CCGS page for further details

\*\*\*\*\*Tissue preferred, but liquid acceptable if tissue insufficient



Choose VA



SHOULDER to SHOULDER  
Every Step of the Way

VA



U.S. Department of Veterans Affairs