THE JANUS RECTAL CANCER TRIAL: A RANDOMIZED PHASE II TRIAL TESTING THE EFFICACY OF TRIPLET VERSUS DOUBLET CHEMOTHERAPY TO ACHIEVE CLINICAL COMPLETE RESPONSE IN PATIENTS WITH LOCALLY ADVANCED RECTAL CANCER

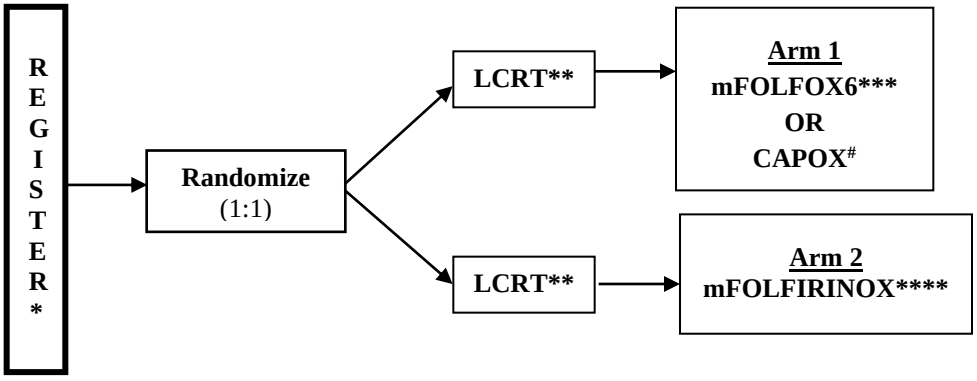
Eligibility Criteria (see Section 3.2)

- Clinical stage II or III rectal adenocarcinoma defined as T4N0, or any T with node positive disease (any T, N+); also T3N0 requiring APR or coloanal anastomosis
- No prior systemic chemotherapy, targeted therapy, or immunotherapy; or radiation therapy administered as treatment for colorectal cancer within the past 5 years
- Not pregnant and not nursing
- Age ≥ 18 years
- ECOG Performance Status 0-1
- No upper rectal tumors (distal margin of tumor > 12 cm from the anal verge)
- No recurrent rectal cancer; prior transanal excision, prior distal sigmoid cancer with a low anastomosis
- No known mismatch repair deficient rectal adenocarcinoma

Required Initial Laboratory Values

ANC	≥ 1500/mm^3
Platelet count:	≥ 100,000/mm^3
Creatinine:	≤ 1.5 x upper limit of normal (ULN) OR
Calc. creatinine clearance:	≥ 50 mL/min
Total bilirubin:	≤ 1.5 x ULN
AST/ALT:	≤ 3 x ULN

Schema



* Patients with locally advanced rectal cancer: <=12cm, T4N0 OR anyT, N1 OR T3N0 that would require APR or coloanal anastomosis

** LCRT = long-course chemoradiation (5 weeks)

***mFOLFOX6 = 8 cycles (1 cycle = 2 weeks)

****mFOLFIRINOX = 8 cycles (1 cycle = 2 weeks)

\# CAPOX = 5 cycles (1 cycle = 3 weeks)

Treatment is to continue for the full course of LCRT and Arm 1 or Arm 2 chemotherapy unless there is a clinical reason to stop. Following neoadjuvant chemotherapy, patients will either proceed to surgery (TME) or watch & wait (WW). Patients will be followed for 8 years or until death, whichever comes first.

Please refer to the full protocol text for a complete description of the eligibility criteria and treatment plan.

LCRT, chemotherapy (as noted in Arm 1 or Arm 2) and surgery will be conducted at the registering institution.

If the Group credited for enrollment is a non-Alliance Group, then other requirements from the credited Group may apply.