## Supplementary

Table S1 Preferred terms included in adverse events of special interest

Adverse event of special interest composite term	MedDRA preferred terms
Neutropenia	Neutropenia, Neutrophil count decreased
Hypertension	Blood pressure increased, Hypertension
Proteinuria	Nephrotic syndrome, Protein urine, Proteinuria, Protein urine present, Urine protein/creatinine ratio abnormal
Hepatic dysfunction	Alanine aminotransferase increased, Aspartate aminotransferase increased, Blood bilirubin increased, Gamma-glutamyl transferase increased, Hepatic function abnormal, Jaundice cholestatic, Liver disorder, Esophageal varices hemorrhage
Leukopenia	Leukopenia, White blood cell count decreased
Febrile neutropenia	Febrile neutropenia
Hemorrhagic events	Conjunctival hemorrhage, Epistaxis, Gingival bleeding, Hematochezia, Hematuria, Esophageal varices hemorrhage, Tumor hemorrhage, Lower gastrointestinal hemorrhage, Post procedural hemorrhage, Hemorrhoidal hemorrhage
Venous thromboembolic events	Pulmonary infarction, Deep vein thrombosis, Embolism, Venous thrombosis limb
Gastrointestinal perforation	Large intestine perforation, Small intestinal perforation
Interstitial lung disease	Interstitial lung disease, Pneumonitis
Arterial thromboembolic events	Cerebral infarction, Pulmonary artery thrombosis
Cardiac failure congestive	Cardiac failure

MedDRA, Medical Dictionary for Regulatory Activities (Version 23.0).

Table S2 Use of FOLFIRI

Parameter	Median (min, max)
Use of levofolinate	
Number of cycles (14 days as 1 cycle)	8 (1, 34)
Duration of treatment (week)	16.4 (2.0, 68.0)
Cumulative dose (mg/kg)	1,165.0 (0.0, 8,125.0)
Dose intensity (mg/kg/week)	80.0 (0.0, 191.3)
Relative dose intensity (%)	80.0 (0.0, 191.3)
Use of fluorouracil	
Number of cycles (14 days as 1 cycle)	8 (1, 34)
Duration of treatment (week)	16.4 (2.0, 68.0)
Cumulative dose (mg/kg)	13,500 (0.0, 108,000.0)
Dose intensity (mg/kg/week)	977.5 (0.0, 2,837.0)
Relative dose intensity (%)	69.8 (0.0, 202.6)
Use of irinotecan	
Number of cycles (14 days as 1 cycle)	8 (1, 34)
Duration of treatment (week)	16.3 (2.0, 68.0)
Cumulative dose (mg/kg)	750.0 (77.0, 5,760.0)
Dose intensity (mg/kg/week)	54.2 (14.4, 143.6)
Relative dose intensity (%)	73.2 (24.0, 114.7)

min, minimum; max, maximum.