

高雄榮民總醫院

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Colon Cancer Clinical Practice Guidelines

Colorectal Cancer Multidisciplinary Team

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Colon Cancer Clinical Practice Guidelines

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■ 本共識依下列參考資料修改版本

□ NCCN Clinical Practical Guidelines in Oncology™ Colon Cancer (Version 1. 2018)

本共識與上一版的差異

上一版	新版
<ol style="list-style-type: none"> 1. Resectable primary colon cancer 2. Adjuvant therapy for stage I-II colon cancer (p.7): 3. Metastatic synchronous adenocarcinoma from large bowel (p.9): 4. Resectable synchronous liver and/or lung metastases only (p.10): 5. Synchronous abdominal/peritoneal metastases (p.12): 6. Chemotherapy for advanced or metastatic disease (p.13-16): 7. Resectable metachronous metastases (p.18) <ol style="list-style-type: none"> a) 8. Chemotherapy regimens for advanced/metastatic disease (p.21-23): 	<ol style="list-style-type: none"> 1. Resectable primary colon cancer (p.6): <ol style="list-style-type: none"> a) Findings: added the category of “Clinical T4b” with the recommendation of “Consider neoadjuvant chemotherapy” followed by “Colectomy with en bloc removal of regional lymph nodes” <p>Locally unresectable or medically inoperable: added the options of “± RT” to “Chemotherapy for Advanced or Metastatic Disease” followed by “Surgery ± IORT or Chemotherapy”</p> <ol style="list-style-type: none"> 2. Adjuvant therapy for stage I-II colon cancer (p.7): <p>Added pathologic stage T3, N0, M0 (MSI high or dMMR) with the recommendation of “no adjuvant therapy.”</p> <p>Surveillance, bullet 3 modified: Chet/abdominal/pelvic CT <i>every 6-12 mo annually</i> for up to 5 y for patients at high risk for recurrence.</p> 3. Metastatic synchronous adenocarcinoma from large bowel (p.9): <p>The following bullet added: Determination of tumor MMR or MSI status (if not previously done)</p> 4. Resectable synchronous liver and/or lung metastases only (p.10): <p>Synchronous or staged colectomy with liver or lung resection (<i>preferred</i>) or <i>local therapy</i>.</p> <p>Footnote “3” added: Resection is preferred over locally ablative procedures (eg, image-guided ablation or SBRT). However, these local techniques can be considered for liver oligometastases</p> 5. Synchronous abdominal/peritoneal metastases (p.12): <p>Footnote “3” added: “Aggressive cytoreductive debulking and/or intraperitoneal chemotherapy are not recommended outside the setting of a clinical trial. If R0 resection can be achieved, surgical resection of isolated peritoneal disease may be considered at experienced centers.”</p> 6. Chemotherapy for advanced or metastatic disease (p.13-16): <p>“Therapy after First Progression”, “Therapy after Second Progression” and “Therapy after Third Progression” were fused as “Subsequent Therapy”</p>

The regimen of trifluridine + tipiracil was added as a subsequent therapy option for patients with disease progression after oxaliplatin- and irinotecan-based chemotherapy.

“Regorafenib (if not given previously) or Trifluridine + tipiracil (if not given previously)” was added in final column as an option

7. [Resectable metachronous metastases](#) (p.18)

b) The treatment option of “resection” modified to “Resection (*preferred*) and/or Local therapy”

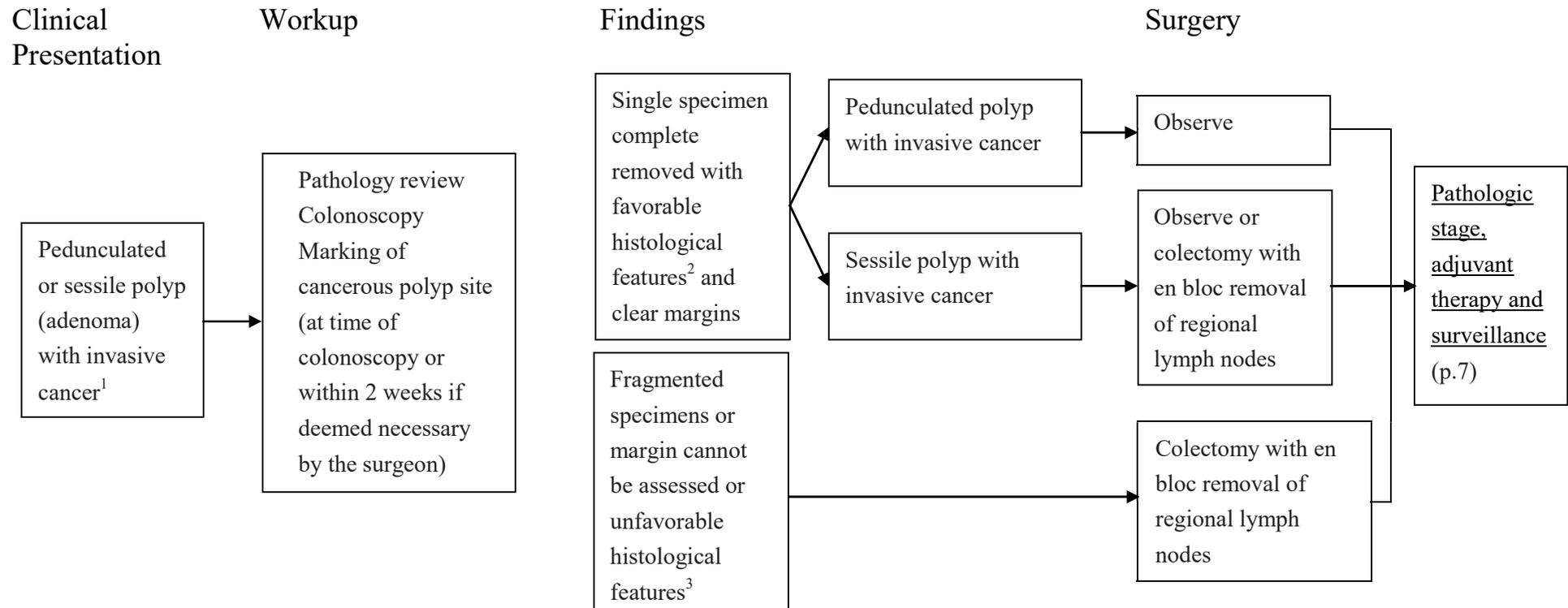
c) Footnote “1” added: “Hepatic artery infusion ± systemic 5-FU/LV is also an option at institutions with experience in both the surgical and medical oncologic aspects of this procedure.”

d) Footnote “2” added: “Resection is preferred over locally ablative procedures (eg, image-guided ablation or SBRT). However, these local techniques can be considered for liver oligometastases.”

8. [Chemotherapy regimens for advanced/metastatic disease](#) (p.21-23):

Regimen added: Trifluridine + tipiracil 35mg/m² up to a Max dose of 80 mg per dose (based on trifluridine component)
PO twice daily 1-5 and 8-12 days repeat every 28 days

Malignant polyp

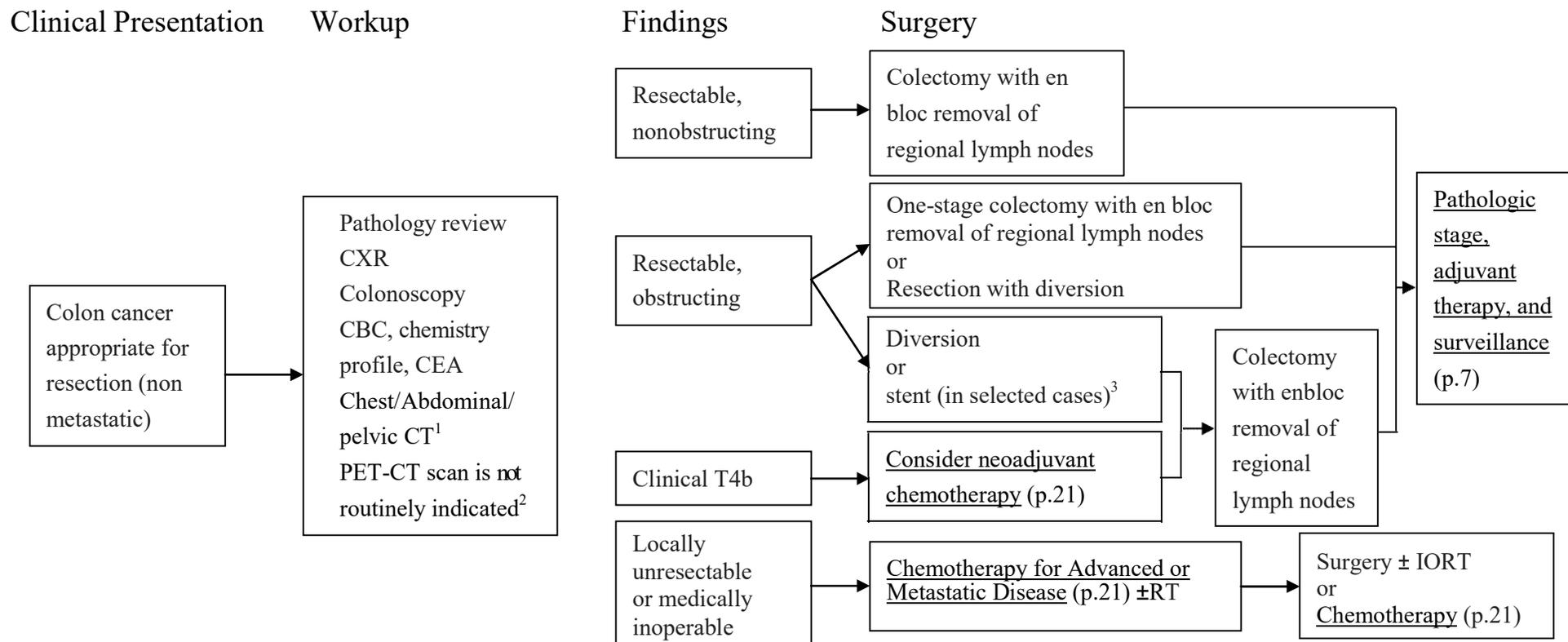


¹A malignant polyp is defined as one with cancer invading through the muscularis mucosae and into the submucosa (pT1). pTis is not considered a “malignant polyp”.

²Favorable histological features: Grade 1 & 2, no angiolymphatic invasion and negative margin of resection

³Unfavorable histological features: Grade 3 & 4, or angiolymphatic invasion, or a “positive” margin (tumour <1mm from the transected margin)

Resectable primary colon cancer

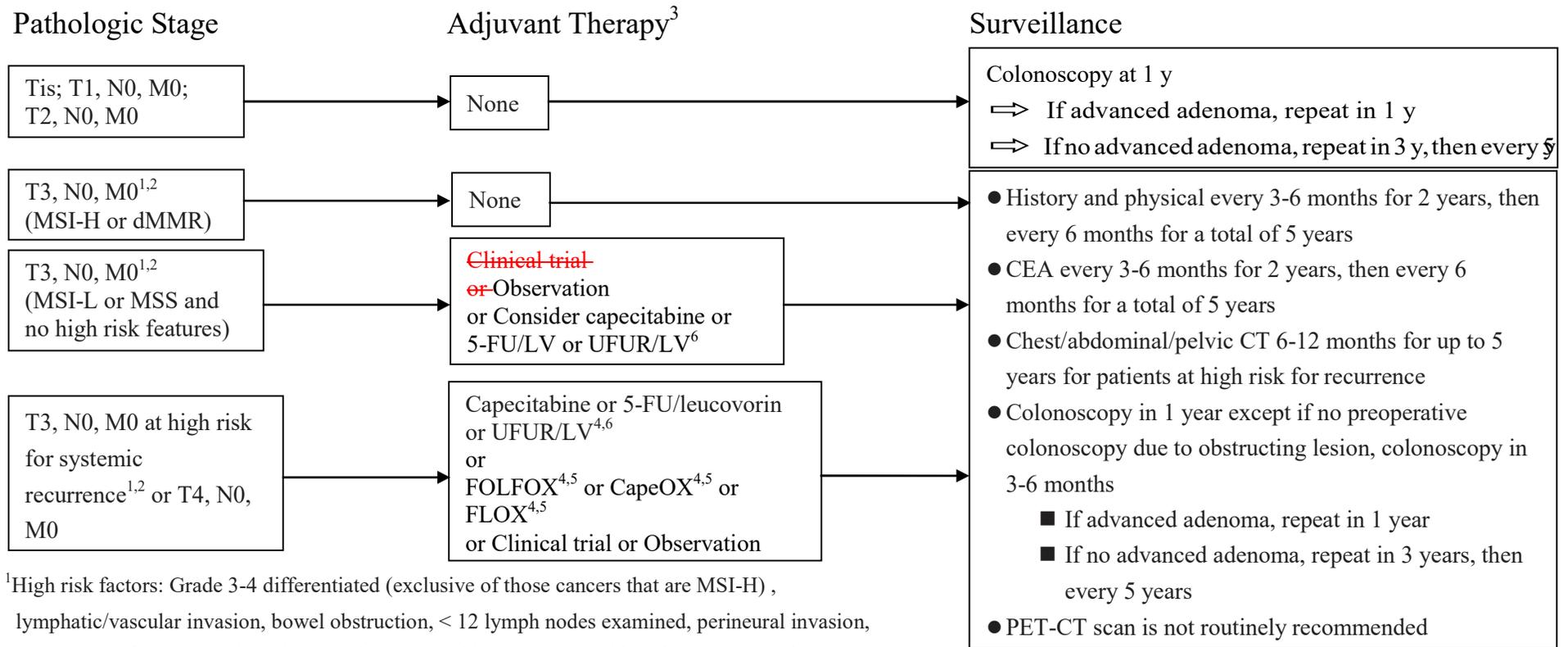


¹CT should be with IV and oral contrast. Consider abd/pelvic MRI with MRI contrast plus a non-contrast chest CT if either CT of abd/pelvis is inadequate or if patient has a contraindication to CT with IV contrast.

²PET-CT does not supplant a contrast-enhanced diagnostic CT scan. PET-CT should only be used to evaluate an equivocal finding on a contrast-enhanced CT scan or in patients with strong contraindications to IV contrast.

³Not available in VGHS now

Adjuvant therapy for stage I-II colon cancer



¹High risk factors: Grade 3-4 differentiated (exclusive of those cancers that are MSI-H), lymphatic/vascular invasion, bowel obstruction, < 12 lymph nodes examined, perineural invasion, localized perforation or close, indeterminate or positive margins. In high-risk stage II patients, there are no data that correlate risk features and selection of chemotherapy.

²Testing for mismatch repair proteins (MMR) should be considered for all patients <70 Y/O or with stage II disease. Stage II MSI-H patients may have a good prognosis and do no benefit from 5-FU adjuvant therapy.

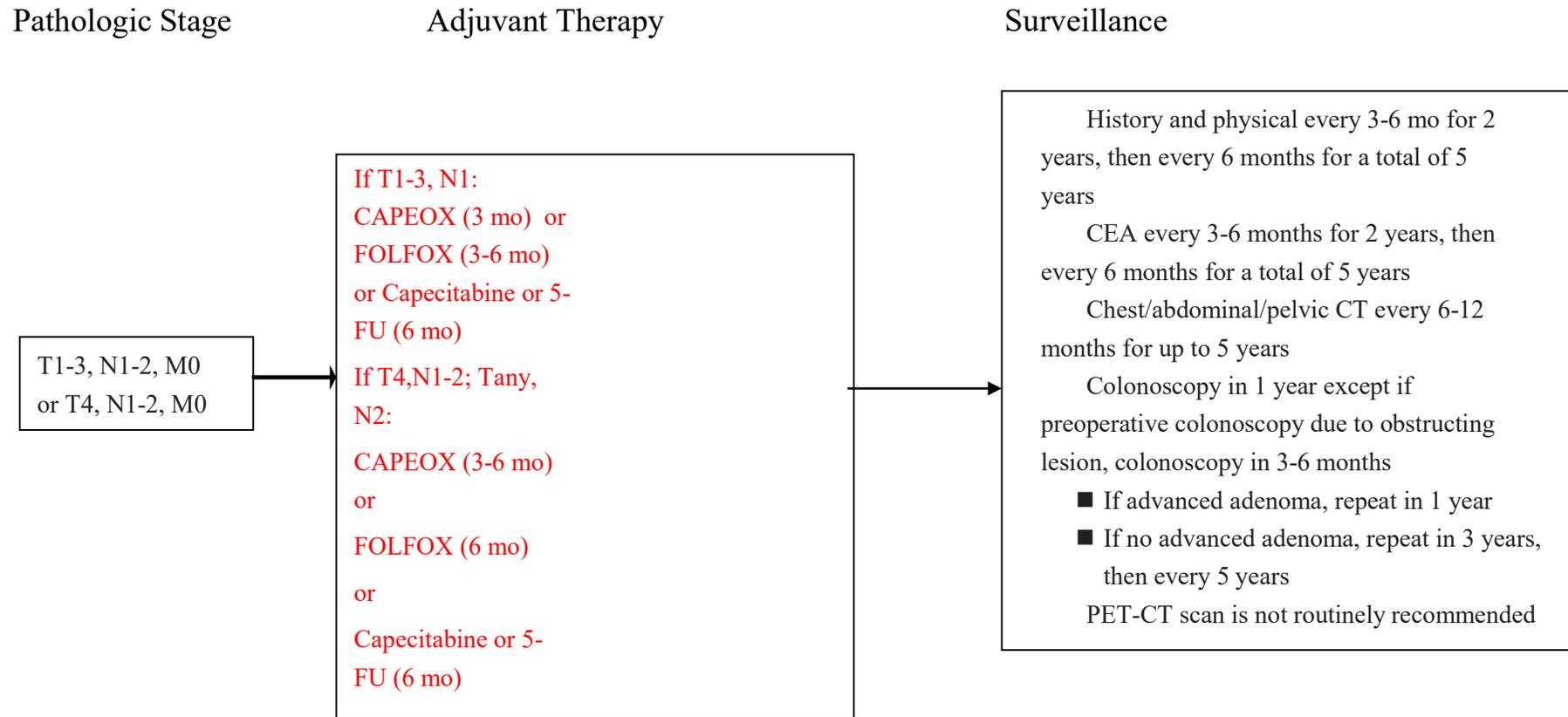
³Bevacizumab, cetuximab, panitumumab, or irinotecan should be not used in the adjuvant setting for stage II or III patients outside the setting of a clinical trial

⁴Consider RT for T4 with penetration to a fixed structure

⁵A benefit for the addition of oxaliplatin to 5-FU/leucovorin in patients age 70 and older has not been proven

⁶Japanese regimen, also see [Chemotherapy Regimens](#)

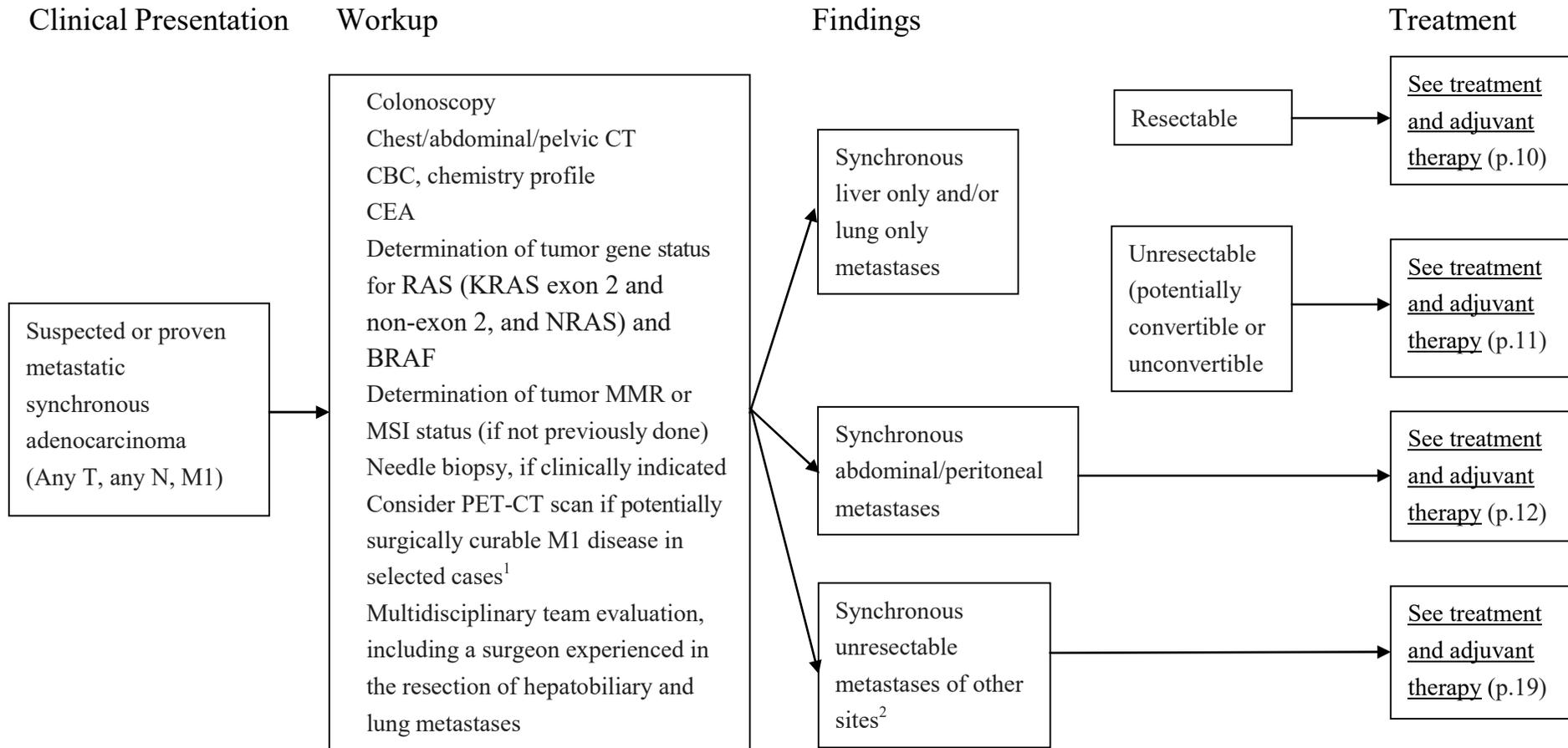
Adjuvant therapy for stage III colon cancer



¹ PLoS One. 2017 Mar 22;12(3):e0174280. Oral tegafur-uracil as metronomic therapy following intravenous FOLFOX for stage III colon cancer.

² Japanese regimen, also see [Chemotherapy Regimens](#)

Metastatic synchronous adenocarcinoma from large bowel



¹Moulton CA, Gu CS, Law CH, et al. Effect of PET before liver resection on surgical management for colorectal adenocarcinoma metastases: a randomized clinical trial. JAMA 2014;311:1863-1869.

²Consider colon resection only if imminent risk of obstruction or significant bleeding.

Resectable synchronous liver and/or lung metastases only

TREATMENT

Resectable synchronous liver and/or lung metastases only

ADJUVANT THERAPY

Resectable metastatic disease
6 months peri-OP treatment preferred²

SURVEILLANCE

If patient stage IV, NED:

- History and physical every 3-6 months for 2 years, then every 6 months for a total of 5 years
- CEA every 3 mo x 2 y, then every 6 months for 3-5 years
- Chest/abdominal/pelvic CT scan every 3-6 months for 2 years, then every 6-12 months up to a total of 5 years
- Colonoscopy in 1 year except if no preoperative colonoscopy due to obstructing lesion, colonoscopy in 3-6 months
 - If advanced adenoma, repeat in 1 years
 - If no advanced adenoma, repeat in 3 years, then every 5 years

Synchronous or staged colectomy with liver or lung resection (preferred) and/or local therapy³

FOLFOX/CapeOx preferred or FOLFIRI ± bevacizumab⁴

Neoadjuvant therapy (for 2-3 months)
FOLFIRI or FOLFOX or CapeOx ± bevacizumab, or **FOLFIRI or FOLFOX ± panitumumab or cetuximab**¹(KRAS/NRAS wild-type [WT] gene only) followed by synchronous or staged colectomy and resection of metastatic disease

Consider observation or shortened course of chemotherapy

Colectomy, followed by chemotherapy (for 2-3 months)
FOLFIRI or FOLFOX or CapeOx ± bevacizumab, or **FOLFIRI or FOLFOX ± panitumumab or cetuximab**¹(KRAS/NRAS WT gene only) and staged resection of metastatic disease

Consider observation or shortened course of chemotherapy

¹There are conflicting data regarding the use of FOLFOX + cetuximab in patients who have potentially resectable liver metastases. ²Total duration of perioperative chemotherapy should not exceed 6 months. ³Resection is preferred over locally ablative procedures (eg, image-guided ablation or SBRT). However, these local techniques can be considered for liver oligometastases

⁴BMC Cancer. 2010 Oct 11;10:545. A randomized two arm phase III study in patients post radical resection of liver metastases of colorectal cancer to investigate bevacizumab in combination with capecitabine plus oxaliplatin (CAPOX) vs CAPOX alone as adjuvant treatment.

Unresectable synchronous liver and/or lung metastases only

TREATMENT

Unresectable synchronous liver and/or lung metastases only

Systemic therapy (FOLFIRI or FOLFOX or CapeOX ± bevacizumab, or FOLFIRI or FOLFOX ± panitumumab or cetuximab [KRAS /NRAS WT gene only] or, FOLFOXIRI ± bevacizumab)
Consider colon resection only if imminent risk of obstruction or significant bleeding

Re-evaluate for conversion to resectable every 2 months if conversion to resectability is a reasonable goal

Converted to resectable

Remains unresectable

Synchronized or staged resection of colon and metastatic cancer

See chemotherapy for advanced or metastatic disease (p.13)

Consider radioembolisation with yttrium-90 resin microspheres for liver limited mets¹

ADJUVANT THERAPY

6 months peri-OP treatment preferred

Active chemotherapy regimen for advanced Disease (Category 2B) or Consider observation or shortened course of chemotherapy

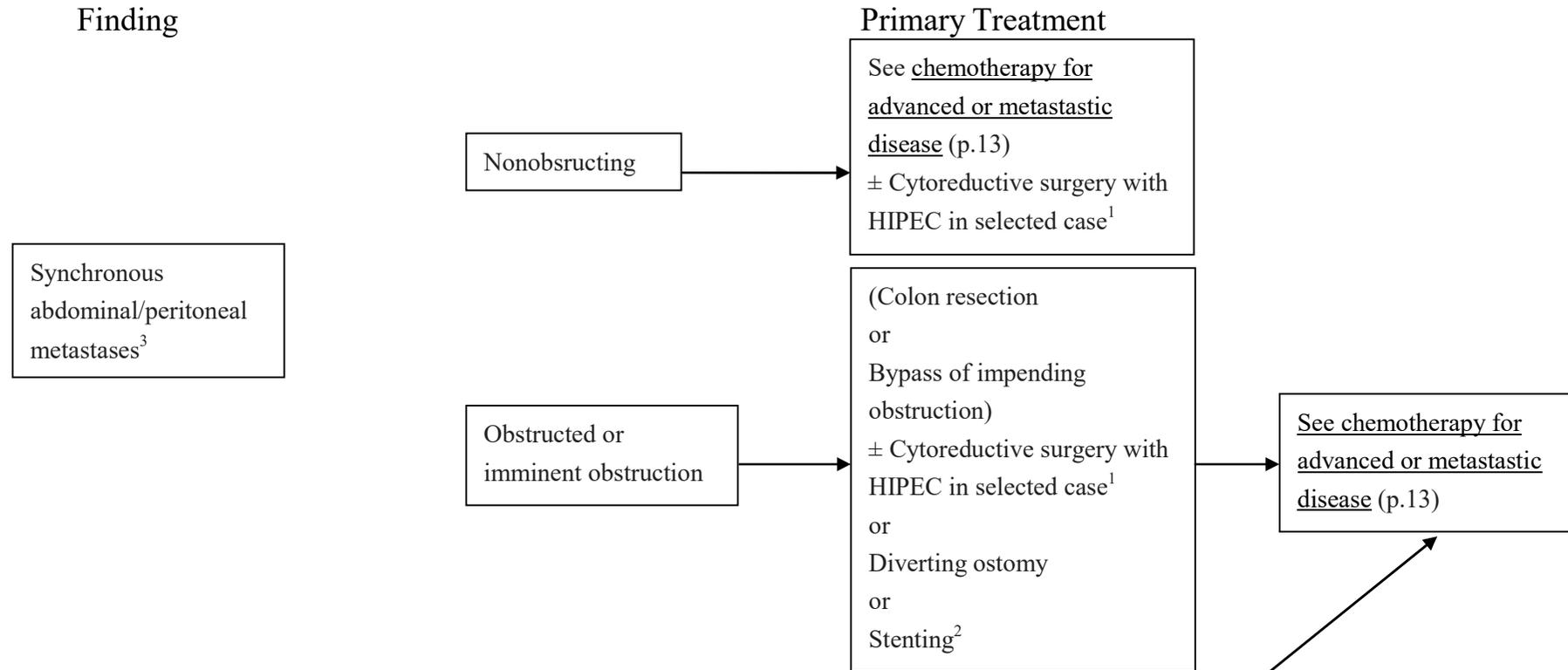
SURVEILLANCE

If patient stage IV, no evidence of disease (NED):

- History and physical every 3-6 months for 2 years, then every 6 months for a total of 5 years
- CEA every 3 months for 2 years, then every 6 months for 3-5 years
- Chest/abdominal/pelvic CT scan every 3-6 months for 2 years, then every 6-12 months up to a total of 5 years
- Colonoscopy in 1 year except if no preoperative colonoscopy due to obstructing lesion, colonoscopy in 3-6 months
 - If advanced adenoma, repeat in 1 year
 - If no advanced adenoma, repeat in 3 years, then every 5 years

¹Not documented in NCCN guideline 2015 v2 but in ESMO guideline 2014(evidence grade IVB). Also refer to reference [9]

Synchronous abdominal/peritoneal metastases

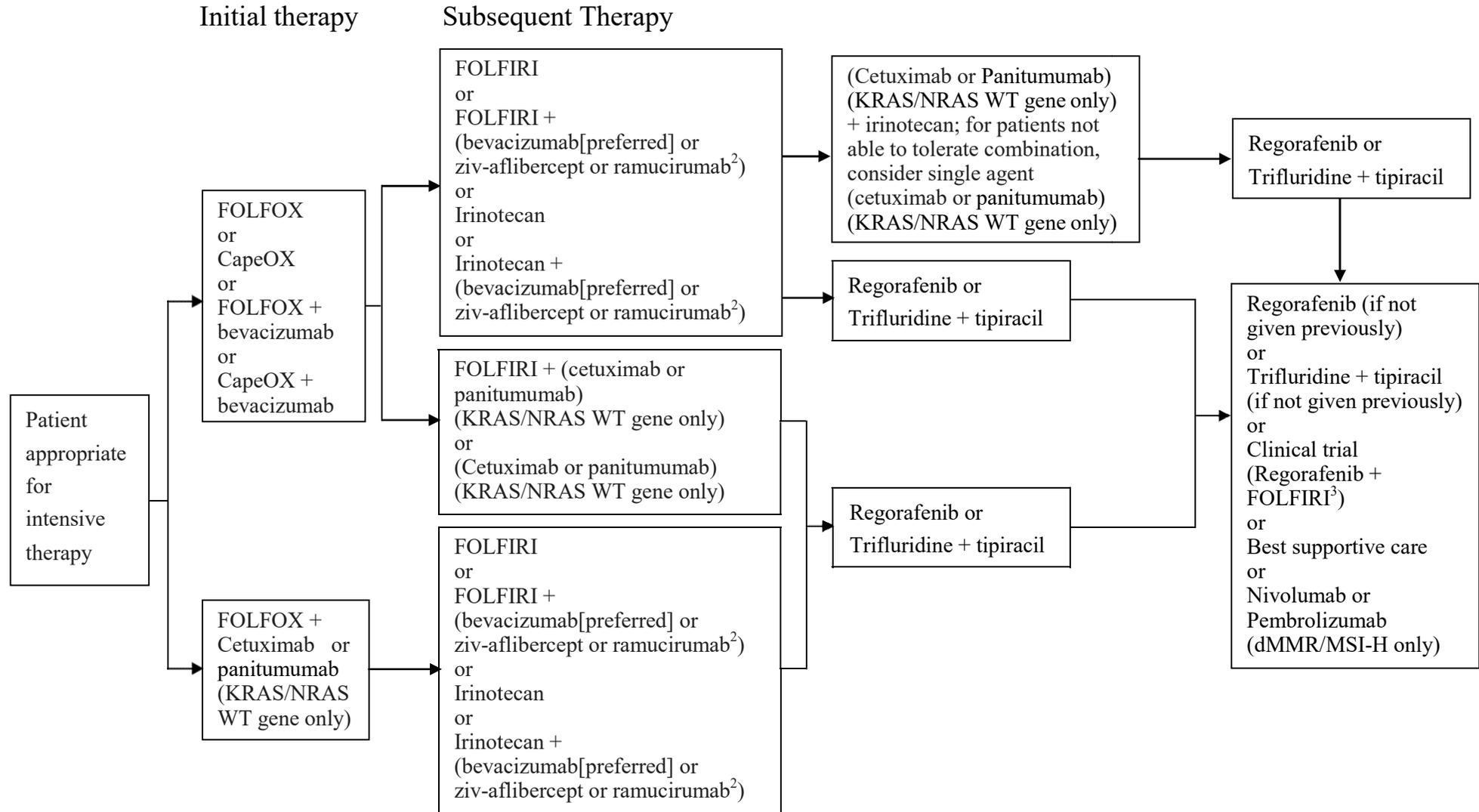


¹HIPEC = Hyperthermic Intraperitoneal Chemotherapy; Not documented in NCCN guideline 2015 v2 but in ESMO guideline 2014 (evidence grade IVB). Also refer to Reference [7], [8]

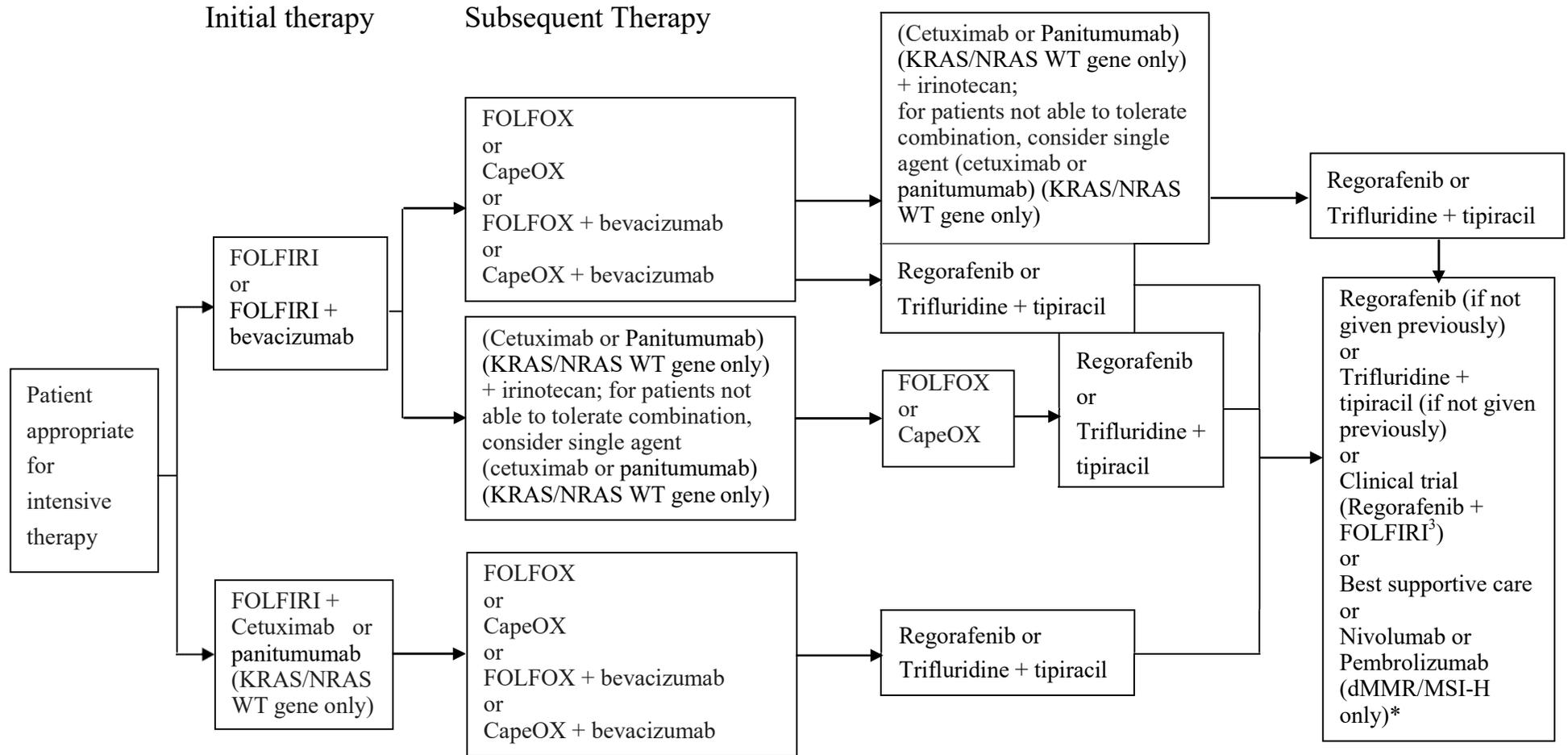
²Not available in VGHS now

³Aggressive cytoreductive debulking and/or intraperitoneal chemotherapy are not recommended outside the setting of a clinical trial. If R0 resection can be achieved, surgical resection of isolated peritoneal disease may be considered at experienced centers.

Chemotherapy for advanced or metastatic disease (1 of 4)



Chemotherapy for advanced or metastatic disease (2 of 4)

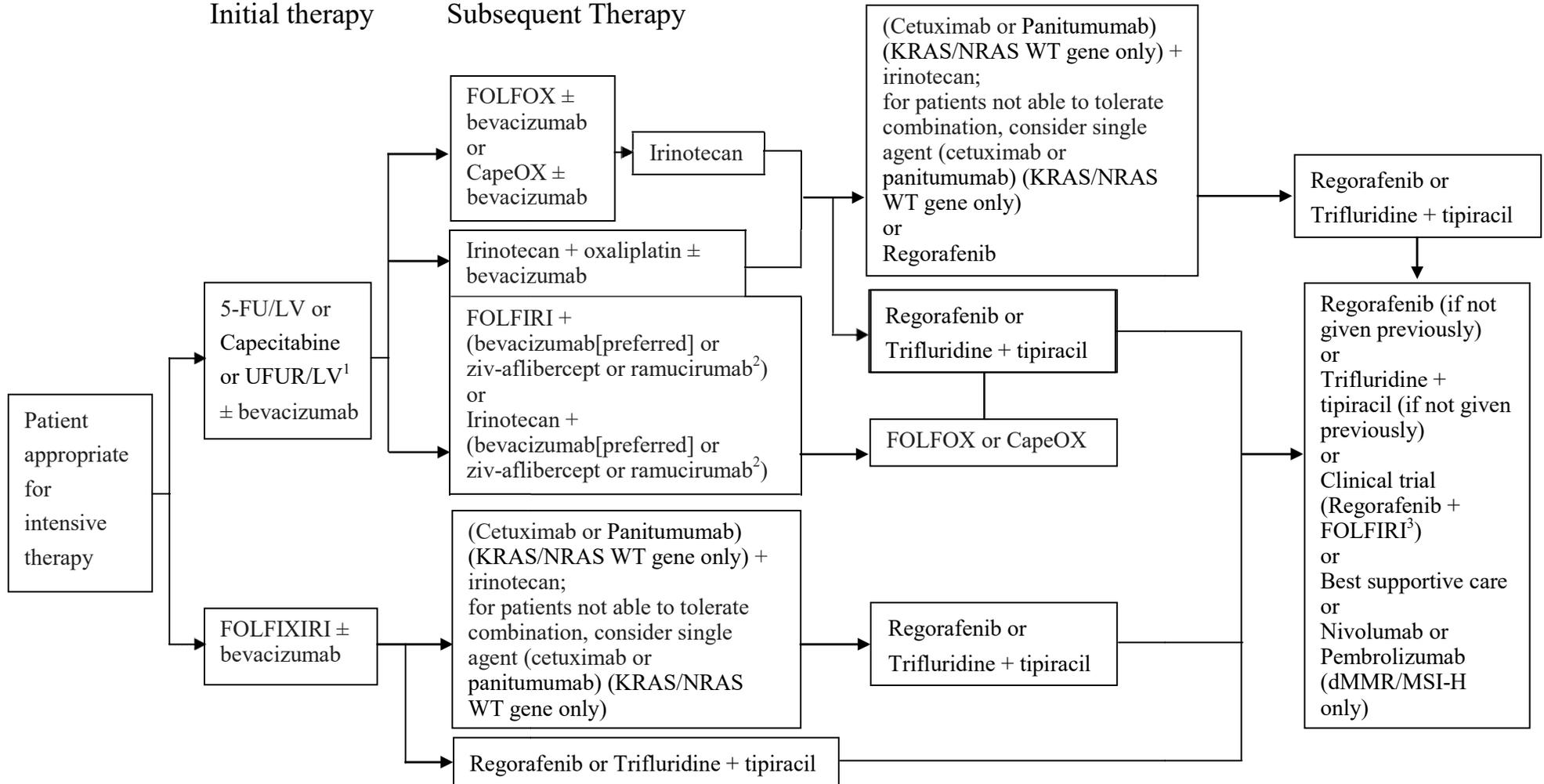


*PD-1 blockade in tumors with mismatch-repair deficiency. N Engl J Med 2015;372:2509-2520. Nivolumab +/- ipilimumab in treatment of patients with metastatic colorectal cancer (mCRC) with and without high microsatellite instability (MSI-H): CheckMate-142 interim results [abstract]. ASCO Meeting Abstracts 2016;34:3501

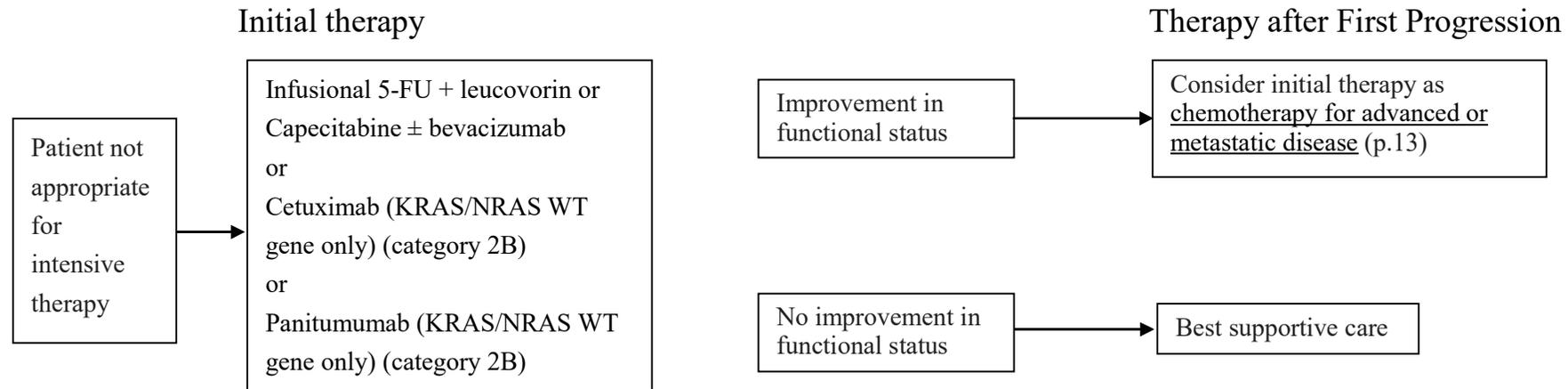
Chemotherapy for advanced or metastatic disease (3 of 4)

Initial therapy

Subsequent Therapy



Chemotherapy for advanced or metastatic disease (4 of 4)



¹Japanese regimen, also see [Chemotherapy Regimens](#)

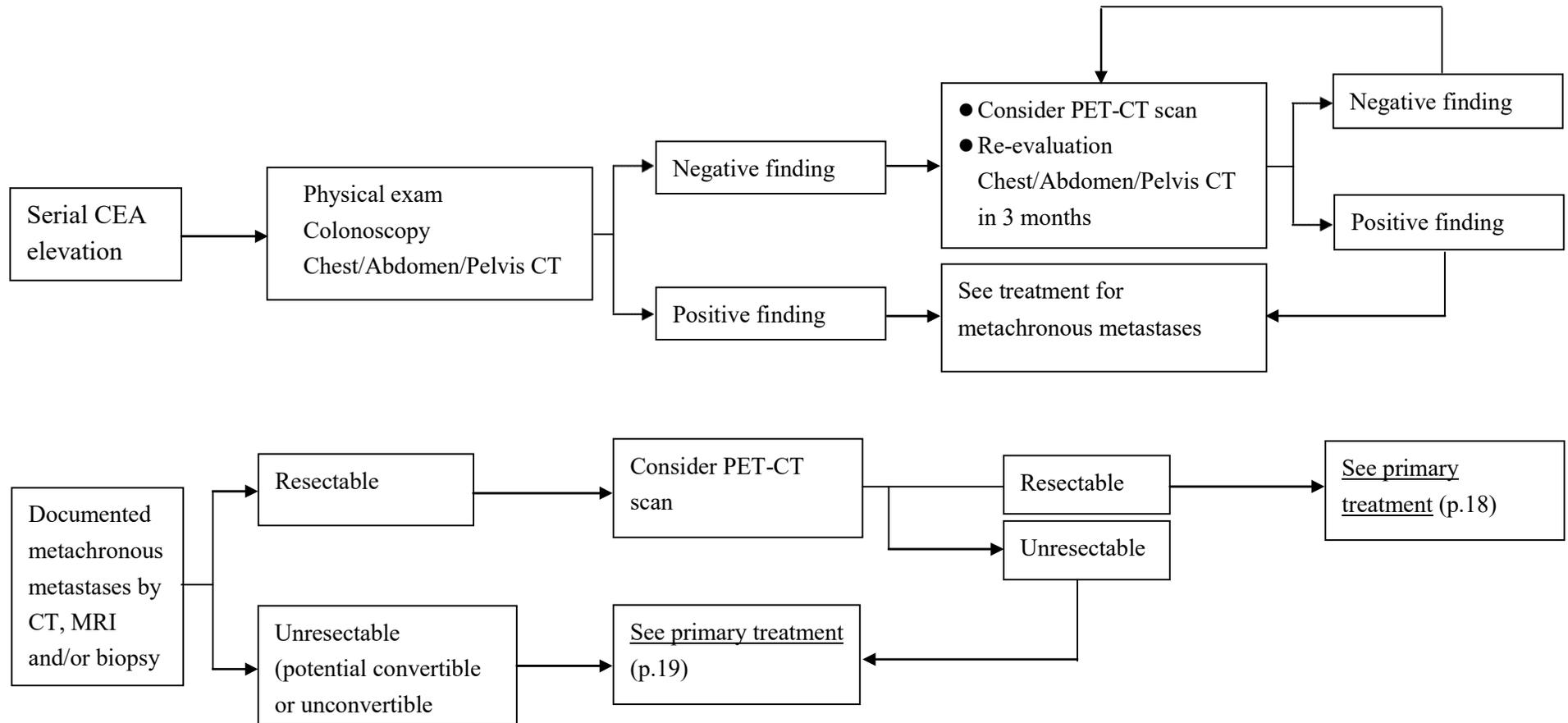
²Not available in routine clinical practice in Taiwan now

³Based on Reference [10], also see footnote "3" in [Chemotherapy Regimens for Advanced/Metastatic Disease \(3 of 3\)](#)

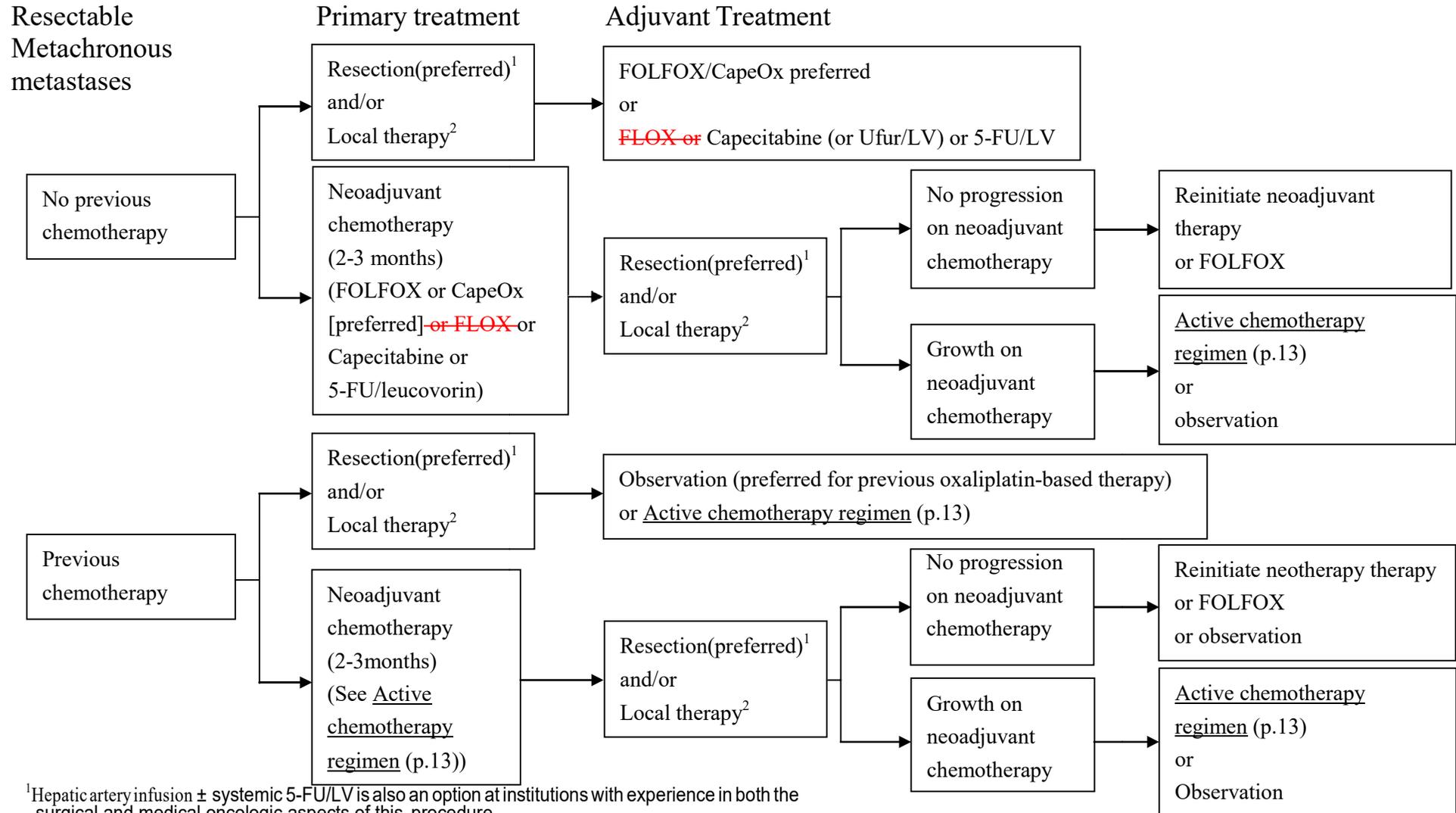
Workup for recurrence

Recurrence

Workup



Resectable metachronous metastases



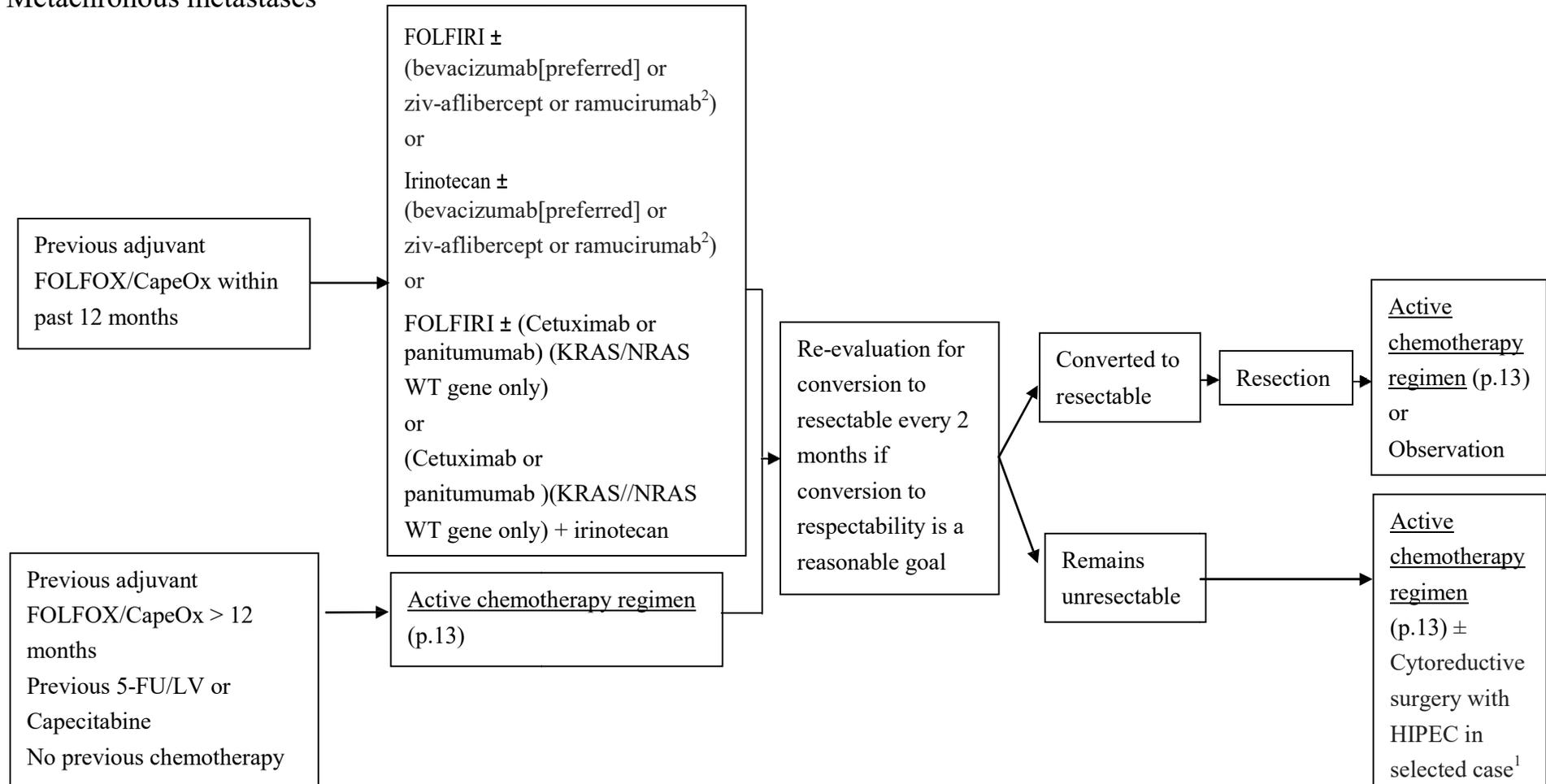
¹Hepatic artery infusion ± systemic 5-FU/LV is also an option at institutions with experience in both the surgical and medical oncologic aspects of this procedure.

²Resection is preferred over locally ablative procedures (eg, image-guided ablation or SBRT). However, these local techniques can be considered for liver oligometastases

Unresectable metachronous metastases

Unresectable
Metachronous metastases

Primary treatment



Unresectable metachronous metastases

¹See footnote “1” in Synchronous abdominal/peritoneal metastases

²Not available in routine practice in Taiwan now

Principles of Chemotherapy

LV Dosage:

Leucovorin 400 mg/m² is the equivalent of levoleucovorin 200 mg/m²

Chemotherapy for Advanced/Metastatic disease

All CRC chemotherapy regimens according to patient's condition and guidelines

NHI regulation:

Bevacizumab combine with Irinotecan base or 5-FU base regimens at the 1st line treatment

Cetuximab combine with Irinotecan or oxaliplatin base regimens at the 1st line & the 3rd line treatment

Panitumumab combine with Irinotecan or oxaliplatin base regimens at the 1st line treatment

Regorafenib at the third/fourth[K-ras wild type] line treatment

Adjuvant Chemotherapy Regimen

Oxaliplatin base (including mFOLFOX6, CapeOX, FLOX)

5-FU base chemotherapy (IV form 5-FU, Capecitabine, Ufur/LV)

NHI regulation:

Oxaliplatin: Stage III colon cancer

Xeloda: Stage III colon cancer, stage IV colorectal cancer

5-FU/LV: High risk stage II, stage III and stage IV colorectal cancer

Ufur/LV: High risk stage II, stage III and stage IV colorectal cancer

Chemotherapy Regimens for Advanced/Metastatic Disease (1 of3)

FOLFOX
<i>mFOLFOX6 (may add with Bevacizumab/Panitumumab/Cetuximab)</i>
Oxaliplatin 85 mg/m ² IV over 2 hours, day 1 Leucovorin 400 mg/m ² IV over 2 hours, day 1 5-FU 400 mg/m ² IV bolus on day 1, then 1200 mg/m ² /day x 2 days (total 2400 mg/m ² over 46–48 hours) IV continuous infusion Repeat every 2 weeks
<i>CapeOX (may add with Bevacizumab)</i>
Oxaliplatin 130 mg/m ² IV over 2 hours, day 1 Capecitabine 850–1000mg/m ² twice daily PO for 14 days Repeat every 3 weeks
FOLFIRI <i>(may add with Bevacizumab/Panitumumab/Cetuximab/Ziv-aflibercept/Ramucirumab)</i>
Irinotecan 180 mg/m ² IV over 30–90 minutes, day 1 Leucovorin* 400 mg/m ² IV infusion to match duration of irinotecan infusion, day 1 5-FU 400 mg/m ² IV bolus day 1, then 1200 mg/m ² /day x 2 days (total 2400 mg/m ² over 46–48 hours) continuous infusion Repeat every 2 weeks
FOLFOXIRI <i>(may add with Bevacizumab)</i>
Irinotecan 165 mg/m ² IV day 1, oxaliplatin 85 mg/m ² day 1, leucovorin 400 mg/m ² day 1, fluorouracil 1600 mg/m ² /day x 2 days (total 3200 mg/m ² over 48 hours) continuous infusion starting on day 1. Repeat every 2 weeks

TARGET THERAPY
Repeat every 2 weeks (unless additional mention)
+ <i>Bevacizumab</i>
Bevacizumab 5 mg/kg IV, day 1 or Bevacizumab 7.5 mg/kg IV, day 1 (for Capecitabine based)
+ <i>Panitumumab (KRAS/NRAS WT gene only)</i>
Panitumumab 6 mg/kg IV over 60 minutes, day 1
+ <i>Cetuximab (KRAS/NRAS WT gene only)</i>
Cetuximab 400 mg/m ² IV over 2 hours first infusion, then 250 mg/m ² IV over 60 minutes weekly or Cetuximab 500 mg/m ² IV over 2 hours, day 1
+ <i>Ziv-aflibercept (FOLFIRI)</i>
Ziv-aflibercept 4 mg/kg IV, day 1
+ <i>Ramucirumab² (FOLFIRI)</i>
Ramucirumab 8mg/kg over 60 minutes, day 1
+ <i>Regorafenib (Single use or with FOLFIRI³)</i>
Regorafenib 160 mg PO daily days 1-21 Repeat every 28 days
<i>Trifluridine + tipiracil²</i>
35mg/m ² up to a Max doas of 80 mg per dose (based on trifluridine component) PO twice daily days 1-5 and 8-12 repeat every 28 days

Chemotherapy Regimens for Advanced/Metastatic Disease (2 of3)

Bolus or infusional 5-FU/leucovorin	Irinotecan based
<i>Roswell Park regimen</i>	<i>IROX</i>
Leucovorin 500 mg/m ² IV over 2 hours, days 1, 8, 15, 22, 29, and 36 5-FU 500 mg/m ² IV bolus 1 hour after start of leucovorin, days 1, 8, 15, 22, 29, and 36 Repeat every 8 weeks	Oxaliplatin 85 mg/m ² IV over 2 hours, followed by irinotecan 200 mg/m ² over 30-90 minutes every 3 weeks
<i>Simplified biweekly infusional 5-FU/LV (sLV5FU2)</i>	<i>Irinotecan (may add with Cetuximab)</i>
Leucovorin 400 mg/m ² IV over 2 hours on day 1, followed by 5-FU bolus 400 mg/m ² and then 1200 mg/m ² /day x 2 days (total 2400 mg/m ² over 46-48 hours) continuous infusion Repeat every 2 weeks	Irinotecan 125 mg/m ² IV over 30-90 minutes, days 1 and 8 Repeat every 3 weeks or Irinotecan 180 mg/m ² IV over 30-90 minutes, day 1 Repeat every 2 weeks or Irinotecan 300-350 mg/m ² IV over 30-90 minutes, day 1 Repeat every 3 weeks
<i>Weekly</i>	
Leucovorin 20 mg/m ² IV over 2 hours on day 1, 5-FU 500 mg/m ² IV bolus injection 1 hour after the start of leucovorin. Repeat weekly. 5-FU 2600 mg/m ² by 24-hour infusion plus leucovorin 500 mg/m ² . Repeat every week (<i>AIO regimen</i> ⁴ : leucovorin 500 mg/m ² in N/S 250ml over 2 hours followed by 5-FU 2600 mg/m ² in N/S 500ml by 24-hour infusion weekly x6 and 2 weeks off, repeat every 8 weeks)	
	Capecitabine (may add with Bevacizumab)
	850–1250 mg/m ² PO twice daily, days 1–14 Repeat every 3 weeks
<i>Mayo Clinic regimen</i> ⁴	Ufur/LV ¹
Leucovorin 20 mg/m ² /day IV over 30 minutes followed by 5-FU IV bolus 425 mg/m ² /day x 5 days. Repeat every 5 weeks	Leucovorin 20-30 mg/m ² + Ufur 300-500 mg/ m ² PO at day 1 to 28 in every 35 days

Chemotherapy Regimens for Advanced/Metastatic Disease (3 of 3)

Modified regimen for CRS@VGHKS
<i>modified mFOLFOX</i>
Oxaliplatin 85-100 mg/ m ² IV over 3 hours on day 1 Leucovorin 200 mg/ m ² IV over 1 hours after Oxaliplatin on day 1 5-FU 2600 mg/m ² IV continuous infusion over 18 hours (start on day 1) Repeat every 2 weeks
<i>modified FOLFIRI</i>
Irinotecan 180 mg/m ² IV over 90 minutes, day 1 Leucovorin 200 mg/m ² IV infusion for 1 hours after irinotecan infusion, day 1 5-FU 2400-3000 mg/m ² continuous infusion over 18 hours (start on day 1) Repeat every 2 weeks
<i>modified AIO regimen</i>
lecovorin 250 mg/m ² in N/S 250ml over 1 hours followed by 5-FU 2600 mg/m ² in N/S 500ml by 18-hour infusion weekly x6 and 2 weeks off, repeat every 8 weeks

¹Japanese regimen, is the equivalent of 5-FU/LV or capecitabine in adjuvant and advanced/metastatic therapy. Also refer to Reference[4], [5] and [6]

²Not available in routine practice in Taiwan now

³As third/fourth line chemotherapy for advanced/metastatic disease, based on reference[10]

⁴At VGHKS

Chemotherapy Regimens for Adjuvant Therapy (1 of 2)

mFOLFOX³	5-FU/leucovorin
Oxaliplatin 85 mg/m ² IV over 2 hours, day 1 Leucovorin 400 mg/m ² IV over 2 hours, day 1 5-FU 400 mg/m ² IV bolus on day 1, then 1200 mg/m ² /day x 2 days (total 2400 mg/m ² over 46–48 hours) IV continuous infusion Repeat every 2 weeks	<i>Rosewell Park regimen (?)</i> Leucovorin 500 mg/m ² given as a 2-hour infusion and repeated weekly x 6. 5-FU 500 mg/m ² given bolus 1 hour after the start of leucovorin and repeated weekly x 6. Every 8 weeks for 4 cycles
FLOX²	<i>Simplified biweekly infusional 5-FU/LV (sLV5FU2)</i>
5-FU 500 mg/m ² IV bolus weekly x 6 + leucovorin 500 mg/m ² IV weekly x 6, each 8-week cycle x 3 with oxaliplatin 85 mg/m ² IV administered on weeks 1, 3, and 5 of each 8-week cycle x 3	Leucovorin 400 mg/m ² IV over 2 hours on day 1, followed by 5-FU bolus 400 mg/m ² and then 1200 mg/m ² /day x 2 days (total 2400 mg/m ² over 46-48 hours) continuous infusion Repeat every 2 weeks
Capecitabine	
1250 mg/m ² PO twice daily, days 1–14 every 3 weeks x 24 wks	
CapeOX	<i>AIO regimen⁴</i>
Oxaliplatin 130 mg/m ² IV over 2 hours, day 1 Capecitabine 850–1000mg/m ² twice daily PO for 14 days Repeat every 3 weeks x 24 weeks	Lecovorin 500 mg/m ² in N/S 250ml over 2 hours followed by 5-FU 2600 mg/m ² in N/S 500ml by 24-hour infusion weekly x6 and 2 weeks off, repeat every 8 weeks
Ufur/LV¹	<i>Mayo Clinic regimen⁴</i>
Leucovorin 20-30 mg/m ² + Ufur 300-500 mg/ m ² PO at day 1 to 28 in every 35 days	Leucovorin 20 mg/m ² /day IV over 30 minutes followed by 5-FU IV bolus 425 mg/m ² /day x 5 days. Repeat every 5 weeks

¹Japanese regimen, is the equivalent of 5-FU/LV or capecitabine in adjuvant and advanced/metastatic therapy. Also refer to Reference[4], [5] and [6]

²FLOX is an alternative to FOLFOX or CapeOx but FOLFOX or CapeOx are preferred

³FOLFOX is reasonable for high-risk or intermediate-risk stage II patients and is not indicated for good- or average-risk patients with stage II colon cancer

⁴At VGHKS

Chemotherapy Regimens for Adjuvant Therapy (2 of 2)

Modified regimen for CRS@VGHKS
<i>modified mFOLFOX</i>
Oxaliplatin 85-100 mg/ m ² IV over 3 hours on day 1 Leucovorin 200 mg/ m ² IV over 1 hours after Oxaliplatin on day 1 5-FU 2600 mg/m ² IV continuous infusion over 18 hours (start on day 1) Repeat every 2 weeks
<i>modified AIO regimen</i>
Lecovorin 250 mg/m ² in N/S 250ml over 1 hours followed by 5-FU 2600 mg/m ² in N/S 500ml by 18-hour infusion weekly x6 and 2 weeks off, repeat every 8 weeks

Definitions for T, N, M	
Primary Tumor (T)	
TX	Primary tumor cannot be assessed
T0	No evidence of primary tumor
Tis	Carcinoma in situ: intraepithelial or invasion of lamina propria ^a
T1	Tumor invades submucosa
T2	Tumor invades muscularis propria
T3	Tumor invades through the muscularis propria into the pericorectal tissues
T4a	Tumor penetrates to the surface of the visceral peritoneum ^b
T4b	Tumor directly invades or is adherent to other organs or structures ^{b,c}
Regional Lymph Nodes (N)	
NX	Regional lymph nodes cannot be assessed
N0	No regional lymph node metastasis
N1a	Metastasis in one regional lymph node
N1b	Metastasis in 2-3 regional lymph nodes
N1c	Tumor deposit(s) in the subserosa, mesentery, or nonperitonealized pericolic or perirectal tissues without regional nodal metastasis
N2a	Metastasis in 4-6 regional lymph nodes
N2b	Metastasis in seven or more regional lymph nodes
Distant Metastasis (M)	
M0	M0 No distant metastasis
M1	M1 Distant metastasis
M1a	Metastasis confined to one organ or site (eg, liver, lung, ovary, onregional node)
M1b	Metastases in more than one organ/site or the peritoneum

7 th AJCC Colorectal cancer staging				Dukes [*]	MAC [*]
Group	T	N	M		
0	Tis	N0	M0	-	-
I	T1	N0	M0	A	A
	T2	N0	M0	A	B1
IIA	T3	N0	M0	B	B2
IIB	T4a	N0	M0	B	B2
IIC	T4b	N0	M0	B	B3
IIIA	T1-2	N1/N1c	M0	C	C1
	T1	N2a	M0	C	C1
IIIB	T3-4a	N1/N1c	M0	C	C2
	T2-3	N2a	M0	C	C1/C2
	T1-2	N2b	M0	C	C1
IIIC	T4a	N2a	M0	C	C2
	T3-4a	N2b	M0	C	C2
	T4b	N1-2	M0	C	C3
IVA	anyT	anyN	M1a	-	-
IVB	anyT	anyN	M1b	-	-

Note: cTNM = clinical classification, pTNM = pathologic classification. Prefix “y” = classification after neoadjuvant pretreatment (eg, ypTNM). Patients who have a complete pathologic response are ypT0N0cM0 that may be similar to Stage Group 0 or I. Prefix “r” = recurred after a disease-free interval (rTNM).

*Dukes B is a composite of better (T3 N0 M0) and worse (T4 N0 M0) prognostic groups, as is Dukes C (Any TN1 M0 and Any T N2 M0). MAC is the modified Astler-Coller classification

癌症藥物停藥準則

1. 根據影像學檢查或臨床依據，針對目前癌症用藥反應效果不良者。
2. 癌症用藥期間，產生藥物不良反應者，或初次發生輕微藥物不良反應後，經調降劑量或處置，仍再次發生藥物不良或更嚴重之反應者。
3. 評估 **adverse effects(AEs)**分級為第三級以上或任何無法承受之併發症者。
4. 評估 Eastern Cooperative Oncology Group(ECOG) Performance Status ≥ 3 者。
5. 經病人意願無法接受及配合持續治療，但經醫師解釋說明後，仍是無法接受癌症用藥或拒絕持續治療者。

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Appendix and Additional Information

1. Dosage of irinotecan in mFOLFIRI + Avstin regimen could be titrated up to 260mg/m² in patient with 6TA/6TA in genotyping of UGT1A1. This is based on the ongoing reseach: **Prospective analysis of UGT1A1 promoter polymorphism for irinotecan dose escalation in metastatic colorectal cancer patients treated with bevacizumab combined with FOLFIRI as the first-line setting** by Dr. Wang