

Systemic Therapy Update

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ULUAVPPPMB

For Health Professionals Who Care for Cancer Patients

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Editor's Choice

New Programs

Effective 01 July 2020, the BC Cancer Provincial Systemic Therapy Program has approved the following new treatment programs. The full details of these programs can be found on the BC Cancer website in the Chemotherapy Protocols section.

Gastrointestinal

First-Line Lenvatinib for Advanced Hepatocellular Carcinoma (UGILEN) — The BC Cancer Gastrointestinal Tumour Group is introducing lenvatinib, a multikinase inhibitor, for the first-line treatment of hepatocellular carcinoma (HCC) that is not amenable to local-regional treatment.¹ Patients may receive lenvatinib or sorafenib (UGISORAF), but not their sequential use, except in the case of intolerance. A

otinib for LUAVPMBM, LUAVPMBM6, LUAVPPMBM, LY: ULYPEM, ULYPEM6

REVISED Protocols, PPPOs and Patient Handouts

NEW Protocols, PPPOs and Patient Handouts

BR: BRAJACTT, BRAJACTTG, BRAVGEMT, BRAVPAM, BRAVTAX, BRAVTRA GI: GIAJFFOX, GIAJRALOX, GIAVCAP, GIDEBTACE, GIFFIRB, UGIFFIRPAN, GIFFOXB, UGIFFOXPAN, GIFIRINOX, GIFOLFIRI, GIFOLFOX, GIGAJFFOX, GIGAVFFOX, GIGAVFFOXT, GIGAVTR, GIGFLODOC, GIGFOLFIRI, UGINETEV, GIPAJFIROX, GIRAJFFOX, GIREGO GO: GOOVBEVG, GOOVBEVLD, GOOVBEVP, GOOVBEVV, GOOVCAG, GOOVCARB, GOOVCATX, GOOVCYCPO, GOOVDOC, GOOVETO, GOOVGEM, GOOVLDOX, GOOVTAX3, GOOVTOP, GOOVVIN GU: GUTAXGEM HN: HNOTLEN LU: LUAVCRIZ, LUAVCRIZF, LUAVPMBM, LUAVPMBM6, ULUAVPPPMB LY: LYNIV, LYNIV4, ULYPRA MY: MYBORPRE, MYBORREL, UMYCARDEX, UMYDARBD, UMYDARLD SC: SCNAUSEA SM: SMAVDT, SMAVI, SMAVNIV, SMAVNIV4, SMAVPEM, SMAVPEM6, SMAVTRA, SMAVVC. SMAVVEM

GI: UGILEN, UGINFOCLAR GO: GOSCPE, GOSCPERT GU: UGUAVPEM,

UGUAVPEM6 LU: LUAVCRIZR, ULUAVPCPMB, ULUAVPGPMB

CAP Requirement Removed:

BR: BRAJPAM, BRAJTTW **GI:** GIAJRALOX, GIDEBTACE **GO:** GOOVBEVG, GOOVBEVLD, GOOVBEVP, GOOVBEVV, GOOVCATB **HN:** HNLADCF **LY:** LYBRENTUX, LYBV, LYIBRU, LYIDELAR, LYMFBEX, LYMIBRU, LYNIV, LYNIV4, LYOBBEND, LYOBCHLOR, LYRITZ, LYVENETOR **SM:** SMAVDAB, SMAVDT, SMAVFIPI, SMAVI, SMAVIPI, SMAVNIV, SMAVNIV4, SMAVPEM, SMAVPEM6, SMAVTRA, SMAVVC, SMAVVEM, SMMCCAVE

Resources and Contact Information

BC Cancer Compassionate Access Program (CAP) approval is required. Patients whose disease progresses on lenvatinib (or sorafenib) may receive regorafenib in the second-line setting (GIREGO).

Approval for this treatment program is based on the phase III REFLECT trial that compared lenvatinib to sorafenib.² The median survival time for lenvatinib was non-inferior to sorafenib (13.6 months vs. 12.3 months, HR 0.92, CI 0.79-1.06). The most common any-grade adverse events associated with lenvatinib were hypertension (42% vs. 30%), diarrhea (39% vs. 46%), decreased appetite (34% vs. 27%), decreased weight (31% vs. 22%), fatigue (30% vs. 25%) and palmar-plantar erythrodysesthesia (27% vs. 52%).

Lenvatinib follows weight-based dosing in the treatment of HCC – in contrast with the fixed-dosing used in thyroid cancer – as summarized in the table below:

Protocols	Lenvatinib Dosing		
	Body weight < 60 kg	Lenvatinib 8 mg daily	
HCC (UGILEN)	Body weight ≥ 60 kg	Lenvatinib 12 mg daily	
Thyroid (UHNOTLEN)	Fixed-dosing	Lenvatinib 24 mg daily	

Octreotide for Non-Functional Neuroendocrine Tumours of the GI Tract (UGINFOCLAR) — The BC Cancer Gastrointestinal Tumour Group is implementing long-acting octreotide (SANDOSTATIN LAR®) for well-differentiated, non-functional neuroendocrine tumours (NETs) of gastrointestinal origin. Until now, only patients with functional NETs have been eligible for therapy with long-acting octreotide (UGIOCTLAR). The phase III randomized, double-blind, placebo-controlled PROMID trial demonstrated improved median time to tumour progression or tumour-related death in patients with both functional and non-functional NETs; significant improvement from 5.9 months to 28.8 months was observed in patients with non-functional, midgut tumours (HR 0.25, 95% CI 0.10-0.59).³ A BC Cancer CAP approval is required. Note that patients with non-functional NETs may receive long-acting octreotide and everolimus (UGINETEV) sequentially, but not in combination.

Lung

First-Line Crizotinib for ROS1-Positive Advanced Non-Small Cell Lung Cancer (LUAVCRIZR) — Crizotinib, a small molecule tyrosine kinase inhibitor, is now approved as first-line therapy for patients with c-ros oncogene 1 (ROS1)-rearranged tumours. ROS1 mutations occur in approximately 1% of patients with non-small cell lung cancer.⁴ Routine ROS1 testing has recently become available, allowing for implementation of this new treatment option. Historically, standard cytotoxic chemotherapy has yielded low response rates and marginal impact on median overall survival in this patient population.

Approval for this program is based on the results of two non-comparative trials, PROFILE 1001 and Ox Onc.^{5,6} Both studies demonstrated clinically meaningful objective response rates (ORR 70% and 71.7%, respectively) and median progression-free survival (mPFS 19.3 months and 15.9 months, respectively). Common any-grade treatment related adverse events (TRAEs) included elevated transaminases, visual disorders, nausea, diarrhea, vomiting, peripheral edema and constipation. The most common grade 3 or 4 TRAEs included neutropenia, elevated transaminases and hypophosphatemia.

SCNAUSEA and Antiemetic Updates

Netupitant-palonosetron, a combination NK₁-5-HT₃ receptor antagonist, was listed as a BC PharmaCare Limited Coverage benefit in November 2019. Although the PharmaCare Collaborative Prescribing Agreement (CPA) for aprepitant was initially replaced by a CPA for netupitant-palonosetron, PharmaCare later reversed its decision to eliminate the aprepitant CPA after consultation with BC Cancer. Thus, both **aprepitant** (plus ondansetron) and **netupitant-palonosetron**, in combination with dexamethasone, are now covered under the PharmaCare CPA for the prevention of acute and delayed chemotherapy-induced nausea and vomiting (CINV) in patients receiving **highly-emetogenic chemotherapy (HEC)**.

Although the PharmaCare funding of netupitant-palonosetron served as a catalyst for updates to BC Cancer protocols and provincial pre-printed orders (PPPOs) in the recent months, netupitant-palonosetron was in fact added to the *BC Cancer Guideline for the Prevention and Treatment of Chemotherapy-Induced Nausea and Vomiting in Adults* (SCNAUSEA) in December 2018, following its inclusion in updated international CINV guidelines. At this time, most BC Cancer protocols and PPPOs have been updated to include aprepitant (plus ondansetron) and netupitant-palonosetron, in combination with dexamethasone, as options for HEC.

Additional SCNAUSEA updates (following the *Prophylactic Antiemetic Regimens* table) include:

- When netupitant-palonosetron is used with anthracycline and cyclophosphamide (AC)-based protocols, omission of day 2 to 4 dexamethasone doses is recommended.
- When netupitant-palonosetron is used, no additional 5-HT₃ receptor antagonist (such as ondansetron) is required, as palonosetron has a long duration of action.
- Aprepitant is the NK₁ receptor antagonist of choice for 3- and 5-day chemotherapy regimens; limited data exist for netupitant-palonosetron. Consult SCNAUSEA or applicable PPPOs for specific dosing details.
- Netupitant is a moderate CYP 3A4 isoenzyme inhibitor, potentially reducing the metabolism of docetaxel and leading to increased docetaxel serum levels. Therefore, aprepitant is considered the NK₁ receptor antagonist of choice in docetaxel-containing regimens. Although docetaxel itself has low emetogenic potential and does not require a NK₁ receptor antagonist or 5-HT₃ receptor antagonist when administered on its own, please consider the following clinical scenarios:
 - When docetaxel is administered in conjunction with HEC (same day of the same cycle), aprepitant-ondansetron is preferred, to avoid an interaction between docetaxel and netupitant (examples of protocols include BRAJDAC, HNAVPD, UHNLADCF and LUAVDC).
 - When docetaxel is administered sequentially with HEC (different cycles), either aprepitantondansetron or netupitant-palonosetron may be selected for the HEC portion of the protocol (examples of protocols include BRAJFECD, BRAJFECDT, BRLAACD and BRLAACDT).
- The moderately-emetogenic chemotherapy (MEC) classification replaced two previous classifications, *high-moderate* and *low-moderate*, in SCNAUSEA in 2018. Some protocols may still contain the outdated classifications. In those instances, the more recent recommendations for MEC should be used to guide antiemetic selection.

References

- 1. Pan-Canadian Oncology Drug Review (pCODR) Expert Review Committee (pERC). Final recommendation for lenvatinib (Lenvima®) for hepatocellular carcinoma. 24 July 2019.
- 2. Kudo M, Finn RS, Qin S, et al. Lenvatinib versus sorafenib in first-line treatment of patients with unresectable hepatocellular carcinoma: a randomized phase 3 non-inferiority trial. *Lancet* 2018;391:1163-1173. <u>https://doi.org/10.1016/S0140-6736(18)30207-1</u>
- 3. Rinke A, Müller H-H, Schade-Brittinger C, et al. Placebo-controlled, double-blind, prospective, randomized study on the effect of octreotide LAR in the control of tumor growth in patients with metastatic neuroendocrine midgut tumors: a report from the PROMID study group. J Clin Oncol 2009;27(28):4656-4663. <u>https://doi.org/10.1200/JCO.2009.22.8510</u>
- 4. Pan-Canadian Oncology Drug Review (pCODR) Expert Review Committee (pERC). Final recommendation for crizotinib (Xalkori®) for ROS-1positive non-small cell lung cancer (NSCLC). 23 May 2019.
- Shaw AT, Riely GJ, Bang Y-J, et al. Crizotinib in ROS1-rearranged advanced non-small-cell lung cancer (NSCLC): updated results, including overall survival, from PROFILE 1001. Ann Oncol 2019;30:1121-1126. <u>https://doi.org/10.1093/announc/mdz131</u>
- 6. Wu Y-L, Yang J C-H, Kim D-W, et al. Phase II study of crizotinib in east Asian patients with ROS1-positive advanced non-small-cell lung cancer. *J Clin Oncol* 2018;36(14):1405-1411. <u>https://doi.org/10.1200/JCO.2017.75.5587</u>

Drug Update

Leucovorin Dosing in Gastrointestinal Protocols

Leucovorin is used to improve the efficacy of 5-fluorouracil (5-FU)-based gastrointestinal therapies. In treatment protocols that employ irinotecan and oxaliplatin, leucovorin is infused concurrently with irinotecan (over 1 hour 30 minutes) or oxaliplatin (over 2 hours), just prior to the administration of 5-FU. Questions about shortening the leucovorin infusion time arise in scenarios where irinotecan or oxaliplatin is omitted. The BC Cancer Gastrointestinal Tumour Group has clarified that there will be no adjustment to the leucovorin infusion when irinotecan or oxaliplatin is omitted from the regimen. Therefore, regardless whether irinotecan or oxaliplatin is given, leucovorin should continue to be administered as 400 mg/m² IV over 1 hour 30 minutes prior to 5-FU in irinotecan-containing protocols, and 400 mg/m² IV over 2 hours prior to 5-FU in oxaliplatin-containing protocols.

Daratumumab Premedications

Daratumumab is used in combination with bortezomib/dexamethasone **(UMYDARBD)** or lenalidomide/dexamethasone **(UMYDARLD)** for the treatment of relapsed and refractory multiple myeloma. Effective 01 July 2020, updates to the *Premedications* section of these protocols have been made; multiple premedications are used for the prevention of daratumumab-associated infusion reactions.

Highlights of revisions to the premedications include:

- Dexamethasone is used both as a therapeutic agent in the treatment of multiple myeloma and for the prevention of daratumumab-associated infusion reactions. Revisions have been made to dexamethasone dosing and days of administration; consult protocol for details based on therapeutic dexamethasone dosage.
- Loratadine, a second-generation antihistamine, has been introduced as an alternative to diphenhydramine. Loratadine is administered prior to each daratumumab infusion, with no need for a repeat dose with longer daratumumab infusions.
- Montelukast, a leukotriene-receptor antagonist, may be omitted after cycle 1 if the patient did not experience infusion reactions with prior cycles.

Note that dexamethasone is dispensed by BC Cancer centres and CON sites, whereas patients must obtain their own supplies of loratadine and montelukast.

Biosimilar Rituximab Coming August 2020

The implementation of biosimilar rituximab will be 01 August 2020. Funding details and updates to applicable rituximab-containing documents will be outlined in the August 2020 Systemic Therapy Update.

Manufacturer Patient Assistance Programs

The listing of patient assistance programs offered by pharmaceutical manufacturers has been updated and can be accessed on the BC Cancer website under Health Professionals > Systemic Therapy > Reimbursement & Forms.

Drug Shortages

The following are updates of drug supply shortages in BC. Full details about new, updated or resolved drug shortages, including recommended treatment alternatives, can be found in the *Briefing Notes* and email communications previously circulated to BC Cancer and the Community Oncology Network (CON).

Updated

Hydroxyurea

(Adapted from BC Cancer Briefing Note Update 12Jun2020)

BC Cancer centres have now received enough supplies to enable dispensing a 4-week supply to patients. Hydroxyurea supplies are expected to stabilize over the next few weeks, which should allow a further increase to dispensing quantities.

Cancer Drug Manual[©]

All BC Cancer Drug Manual[©] documents can be accessed from the <u>Cancer Drug Manual[©]</u> home page on the BC Cancer website.

Revised Documents

Highlights of key changes are listed below:

Atezolizumab Monograph

Dosage Guidelines: added new 2-weekly and 4-weekly regimens

Avelumab Monograph

Uses: added new indications Dosage Guidelines: added new fixed-dose regimen

Cyclophosphamide Chemotherapy Preparation and Stability Chart

Product and *Product Stability* columns: updated suggested volumes and final product stability following product review

Docetaxel Monograph

Interactions: added netupitant interaction to table

Mitomycin Monograph

Solution Preparation and Compatibility: added information relating to NS as an alternate diluent for reconstitution

Olaparib Monograph

Dosage Guidelines: updated hepatic dosing

Provincial Systemic Therapy Program

Financial Support Drug Program (FSDP) Update

The Financial Support Drug Program (FSDP) has been jointly operated by BC Cancer and the Canadian Cancer Society (CCS) to provide symptom control medications for oncology patients. CCS was providing the financial screening that enabled patients to be enrolled in the program, however, this service closed as of 01 June 2020. Until further notice, patients with current approval will continue to receive funding through the program; however, no new patients will be enrolled. Any patient whose approval is expiring or new patients seeking assistance with funding for supportive care medications are asked to contact their regional BC Cancer Patient and Family Counseling department to explore other options:

www.bccancer.bc.ca/our-services/services/supportive-care/patient-family-counselling.

Research

Immune Checkpoint Inhibition in Renal Transplant Recipients

A Precision-Medicine-Guided Approach to Multidisciplinary Care

Kidney transplant recipients are at elevated risk of cancer compared with the general population. This increased risk is largely due to the immunosuppressive effects of the anti-rejection medications that are required to prevent transplant graft rejection. The risk of certain cancers, including melanoma, lung cancer and kidney cancer, is especially increased among kidney transplant recipients.

For transplant patients receiving systemic therapy for the management of cancer, clinicians often reduce the intensity of maintenance anti-rejection medications to help control the cancer; however, this may increase the risk of graft rejection and failure. Immune checkpoint inhibitors, which may be used in the management of cancer, may also contribute to graft rejection; checkpoint inhibitor use in organ transplant recipients has resulted in rejection in 37% to 50% of cases.

Researchers from the University of Washington, the renal transplant group in Vancouver, and BC Cancer are collaborating on an upcoming study to optimize the care of kidney transplant recipients who are eligible for checkpoint inhibitor therapy for the management of cancer. The study aims to characterize the risk of graft rejection in adult kidney transplant recipients, and to perform screening for rejection and surveillance for cancer recurrence in checkpoint-inhibitor-treated kidney transplant recipients.

Recent advances in the use of cell-free DNA have enabled earlier diagnoses of graft rejection and cancer recurrence; the precision-medicine portion of the study will be guided by the study's principal investigator, Dr. Chris Blosser (nephrologist, University of Washington) and Dr. James Lan (transplant nephrologist, Vancouver General Hospital). An additional goal of the study is to streamline the referral process to the renal transplant group (represented by Dr. Lan and Dr. John Gill, St. Paul's Hospital) for checkpoint inhibitor-eligible kidney transplant recipients. Creation of a multidisciplinary framework will be investigated by Dr. Sanjay Rao (medical oncologist, BC Cancer – Kelowna); multidisciplinary collaboration will ensure that patients receive optimal testing, treatment and follow-up, to improve graft-and cancer-related outcomes.

Submitted by Drs. Blosser, Lan and Rao

Contact Dr. Sanjay Rao at <u>SRao@bccancer.bc.ca</u>

Systemic Therapy Update Editorial Board

Membership Update

The Systemic Therapy Update Editorial Board would like to bid farewell to **Dr. Sally Waignein** (Provincial Pharmacy Education Coordinator, BC Cancer) as she leaves BC Cancer. Dr. Waignein joined the ST Update Editorial Board in 2011, serving as editor from 2012 to 2019, and continued to support the ST Update in the assistant editor role. The ST Update has benefited enormously from her oversight and wishes her all the best in her next chapter.

Benefit Drug List

New Programs

Effective 01 July 2020, the following new treatment programs have been added to the BC Cancer <u>Benefit</u> <u>Drug List</u>:

Protocol Title	Protocol Code	Benefit Status
First-Line Therapy of Advanced Hepatocellular Carcinoma using Lenvatinib	UGILEN	Restricted
Management of Non-Functional Neuroendocrine Tumours of the GI Tract using Octreotide (SANDOSTATIN LAR[®])	UGINFOCLAR	Restricted
First-Line Treatment of ROS1-Positive Advanced Non-Small Cell Lung Cancer with Crizotinib	LUAVCRIZR	Class I

Revised Programs

Effective 01 July 2020, the following treatment program has been revised on the BC Cancer <u>Benefit</u> <u>Drug List</u>:

Protocol Title	Protocol Code	Benefit Status
Management of Non-Functional Neuroendocrine Tumours of the GI Tract using Octreotide	GINOS	Class I (expanded Eligibility)

The following programs have been transferred to Class I status on the BC Cancer Benefit Drug List:

Protocol Title	Protocol Code	Benefit Status
Adjuvant Therapy in Postmenopausal Women using Pamidronate	BRAJPAM	Class I (previously Restricted)
Adjuvant Therapy using Weekly Paclitaxel and Trastuzumab	BRAJTTW	Class I (previously Restricted)

Benefit Drug List

Adjuvant Chemotherapy for Node-Positive Colon Cancer using Oxaliplatin and Raltitrexed in Patients Intolerant to Fluorouracil or Capecitabine	GIAJRALOX	Class I (previously Restricted)
Transarterial Chemoembolization (TACE) of Hepatocellular Carcinoma using Drug-Eluting Bead (DEB) Loaded with Doxorubicin	GIDEBTACE	Class I (previously Restricted)
Treatment of Platinum-Resistant Ovarian Cancer with Bevacizumab and Gemcitabine	GOOVBEVG	Class I (previously Restricted)
Treatment of Platinum-Resistant Ovarian Cancer with Bevacizumab and Doxorubicin Pegylated Liposomal (CAELYX®)	GOOVBEVLD	Class I (previously Restricted)
Treatment of Platinum-Resistant Ovarian Cancer with Bevacizumab and Paclitaxel	GOOVBEVP	Class I (previously Restricted)
Treatment of Platinum-Resistant Ovarian Cancer with Bevacizumab and Vinorelbine	GOOVBEVV	Class I (previously Restricted)
Primary Treatment of Invasive Ovarian with High-Risk of Relapse using Bevacizumab, Carboplatin and Paclitaxel	GOOVCATB	Class I (previously Restricted)
Locally Advanced Squamous Cell Carcinoma with Docetaxel, Cisplatin and Infusional Fluorouracil	HNLADCF	Class I (previously Restricted)
Treatment of Hodgkin Lymphoma and Anaplastic Large Cell Lymphoma with Brentuximab Vedotin	LYBRENTUX	Class I (previously Restricted)
Consolidation Therapy Post-ASCT for Hodgkin Lymphoma using Brentuximab Vedotin	LYBV	Class I (previously Restricted)
Treatment of Relapsed/Refractory Chronic Lymphocytic Leukemia using Ibrutinib	LYIBRU	Class I (previously Restricted)
Treatment of Relapsed/Refractory Chronic Lymphocytic Leukemia using Idelalisib and Rituximab	LYIDELAR	Class I (previously Restricted)
Treatment of Cutaneous T-Cell Lymphoma (Mycosis Fungoides/Sézary Syndrome) with Bexarotene	LYMFBEX	Class I (previously Restricted)
Treatment of Relapsed/Refractory Mantle-Cell Lymphoma using Ibrutinib	LYMIBRU	Class I (previously Restricted)
Treatment of Relapsed or Refractory Hodgkin Lymphoma using Nivolumab	LYNIV	Class I (previously Restricted)
Treatment of Relapsed or Refractory Hodgkin Lymphoma using 4-Weekly Nivolumab	LYNIV4	Class I (previously Restricted)
Treatment of Rituximab-Refractory Follicular Lymphoma with Obinutuzumab in Combination with Bendamustine	LYOBBEND	Class I (previously Restricted)
Treatment of Previously-Untreated Chronic Lymphocytic Leukemia with Obinutuzumab and Chlorambucil	LYOBCHLOR	Class I (previously Restricted)
Treatment of Lymphoma using Radioimmunotherapy: Rituximab -Priming for Ibritumomab ⁹⁰ Y (ZEVALIN®)	LYRITZ	Class I (previously Restricted)

Benefit Drug List

Treatment of Relapsed/Refractory Chronic Lymphocytic Leukemia using Venetoclax and Rituximab	LYVENETOR	Class I (previously Restricted)
Treatment of BRAF-Positive Unresectable or Metastatic Melanoma using Dabrafenib	SMAVDAB	Class I (previously Restricted)
Treatment of BRAF-Positive Unresectable or Metastatic Melanoma using Dabrafenib and Trametinib	SMAVDT	Class I (previously Restricted)
First-Line Treatment of Unresectable or Metastatic Melanoma using Ipilimumab	SMAVFIPI	Class I (previously Restricted)
Treatment of Advanced C-Kit-Positive Melanoma using Imatinib	SMAVI	Class I (previously Restricted)
Treatment of Unresectable or Metastatic Melanoma using Ipilimumab	SMAVIPI	Class I (previously Restricted)
Treatment of Unresectable or Metastatic Melanoma using Nivolumab	SMAVNIV	Class I (previously Restricted)
Treatment of Unresectable or Metastatic Melanoma using 4-Weekly Nivolumab	SMAVNIV4	Class I (previously Restricted)
Treatment of Unresectable or Metastatic Melanoma using Pembrolizumab	SMAVPEM	Class I (previously Restricted)
Treatment of Unresectable or Metastatic Melanoma using 6-Weekly Pembrolizumab	SMAVPEM6	Class I (previously Restricted)
Treatment of BRAF-Positive Unresectable or Metastatic Melanoma using Trametinib	SMAVTRA	Class I (previously Restricted)
Treatment of BRAF-Positive Unresectable or Metastatic Melanoma using Vemurafenib and Cobimetinib	SMAVVC	Class I (previously Restricted)
Treatment of BRAF-Positive Unresectable or Metastatic Melanoma using Vemurafenib	SMAVVEM	Class I (previously Restricted)
Second-Line Treatment of Recurrent or Metastatic Merkel Cell Carcinoma using Avelumab	SMMCCAVE	Class I (previously Restricted)

Highlights of New & Revised Protocols, PPPOs and Patient Handouts

BC Cancer Protocol Summaries, Provincial Pre-Printed Orders (PPPOs) and Patient Handouts are revised periodically. New, revised or deleted protocols, PPPOs and patient handouts for this month are listed below, with document revisions indicated in the respective columns. Protocol codes for treatment requiring BC Cancer Compassionate Access Program approval are prefixed with the letter **U**.

NEW Protocols, PPPOs and Patient Handouts (new documents checked 🗹)				
Code	Protocol Title	Protocol	РРРО	Handout
UGILEN	First-Line Therapy of Advanced Hepatocellular Carcinoma using Lenvatinib	\checkmark	V	
UGINFOCLAR	Management of Non-Functional Neuroendocrine Tumours of the GI Tract using Octreotide (SANDOSTATIN LAR [®])	\checkmark	V	\checkmark
GOSCPE	Treatment of Small Cell Gynecologic Cancer with Cisplatin and Etoposide			V
GOSCPERT	Treatment of Small Cell Gynecologic Cancer using Cisplatin and Etoposide with Radiation Therapy			\checkmark
UGUAVPEM	Treatment of Locally Advanced or Metastatic Urothelial Carcinoma using Pembrolizumab			V
UGUAVPEM6	Treatment of Locally Advanced or Metastatic Urothelial Carcinoma using 6-Weekly Pembrolizumab			
LUAVCRIZR	First-Line Treatment of ROS1-Positive Advanced Non-Small Cell Lung Cancer with Crizotinib	\checkmark	$\overline{\checkmark}$	\checkmark
ULUAVPCPMB	First-Line Treatment of Advanced Squamous Non-Small Cell Lung Cancer with Paclitaxel, Carboplatin and Pembrolizumab			\checkmark
ULUAVPGPMB	First-Line Treatment of Advanced Squamous Non-Small Cell Lung Cancer with Platinum, Gemcitabine and Pembrolizumab			Carboplatin and cisplatin options
LUAVPMBM	Maintenance Therapy of Advanced Non-Small Cell Lung Cancer with Pembrolizumab			V
LUAVPMBM6	Maintenance Therapy of Advanced Non-Small Cell Lung Cancer with 6-Weekly Pembrolizumab			\checkmark
LUAVPPMBM	Maintenance Therapy of Advanced Non- Squamous Non-Small Cell Lung Cancer with Pemetrexed and Pembrolizumab			V

NEW Protocols, PPPOs and Patient Handouts (new documents checked 🗹)				
Code	Protocol Title	Protocol	РРРО	Handout
ULUAVPPPMB	First-Line Treatment of Advanced Non- Squamous Non-Small Cell Lung Cancer with Platinum, Pemetrexed and Pembrolizumab			Carboplatin and cisplatin options
ULYPEM	Treatment of Relapsed or Refractory Hodgkin Lymphoma using Pembrolizumab			V
ULYPEM6	Treatment of Relapsed or Refractory Hodgkin Lymphoma using 6-Weekly Pembrolizumab			V

REVISED Protocols, PPPOs and Patient Handouts (revisions in respective columns)

				,	
Code	Protocol Title	Protocol	РРРО	Handout	
BR Breast	BR Breast				
BRAJACTT	Neoadjuvant or Adjuvant Therapy for Breast Cancer using Doxorubicin and Cyclophosphamide followed by Paclitaxel and Trastuzumab	Cyclophosphamide and paclitaxel bag size clarified			
BRAJACTTG	Neoadjuvant or Adjuvant Therapy for Breast Cancer using Dose-Dense Therapy: Doxorubicin and Cyclophosphamide Followed by Paclitaxel and Trastuzumab	Cyclophosphamide and paclitaxel bag size clarified			
BRAVGEMT	Palliative Therapy for Metastatic Breast Cancer using Gemcitabine and Paclitaxel	Paclitaxel bag size clarified			
BRAVPAM	Prevention of Skeletal-Related Events Secondary to Breast Cancer Metastases using Pamidronate	Renal impairment dosing clarified			
BRAVTAX	Palliative Therapy for Metastatic Breast Cancer using Paclitaxel	Paclitaxel bag size clarified			
BRAVTRAP	Palliative Therapy for Metastatic Breast Cancer using Trastuzumab and Paclitaxel as First-Line Treatment for Advanced Breast Cancer	Paclitaxel bag size clarified			
GI Gastrointe	estinal				
GIAJFFOX	Adjuvant Combination Chemotherapy for Stage III and Stage IIB Colon Cancer using Oxaliplatin, Fluorouracil and Leucovorin	Leucovorin dosing (see Drug Update) and oxaliplatin bag size clarified			
GIAJRALOX	Adjuvant Combination Chemotherapy for Node-Positive Colon Cancer using Oxaliplatin and Raltitrexed in Patients Intolerant to Fluorouracil or Capecitabine	Oxaliplatin bag size clarified	Oxaliplatin bag size clarified		

REVISED Protocols, PPPOs and Patient Handouts (revisions in respective columns)

Code	Protocol Title	Protocol	РРРО	Handout
GIAVCAP	Palliative Therapy of Advanced Colorectal Cancer using Capecitabine		CAP statement deleted	
GIDEBTACE	Transarterial Chemoembolization (TACE) of Hepatocellular Carcinoma using Drug-Eluting Bead (DEB) Loaded with Doxorubicin	Institution name and Tests revised		
GIFFIRB	Palliative Combination Chemotherapy for Metastatic Colorectal Cancer using Irinotecan, Fluorouracil, Leucovorin and Bevacizumab	Leucovorin dosing clarified (see Drug Update)		
UGIFFIRPAN	Palliative Combination Chemotherapy for Metastatic Colorectal Cancer using Irinotecan, Fluorouracil, Leucovorin and Panitumumab	Leucovorin dosing clarified (see Drug Update)		
GIFFOXB	Palliative Combination Chemotherapy for Metastatic Colorectal Cancer using Oxaliplatin, Fluorouracil, Leucovorin and Bevacizumab	Leucovorin dosing (see Drug Update) and oxaliplatin bag size clarified		
UGIFFOXPAN	Palliative Combination Chemotherapy for Metastatic Colorectal Cancer using Oxaliplatin, Fluorouracil, Leucovorin and Panitumumab	Leucovorin dosing (see Drug Update) and oxaliplatin bag size clarified		
GIFIRINOX	Palliative Combination Chemotherapy for Advanced Pancreatic Adenocarcinoma using Irinotecan, Oxaliplatin, Fluorouracil and Leucovorin	Leucovorin dosing (see Drug Update) and oxaliplatin bag size clarified		
GIFOLFIRI	Palliative Combination Chemotherapy for Metastatic Colorectal Cancer using Irinotecan, Fluorouracil and Leucovorin	Leucovorin dosing clarified (see Drug Update)		
GIFOLFOX	Palliative Combination Chemotherapy for Metastatic Colorectal Cancer using Oxaliplatin, Fluorouracil and Leucovorin	Leucovorin dosing (see Drug Update) and oxaliplatin bag size clarified		
GIGAJFFOX	Adjuvant Chemotherapy of Gastric Cancer Patients with D2 Resection (Node-Negative) or Ineligible for Adjuvant Chemoradiation using Oxaliplatin, Fluorouracil and Leucovorin	Leucovorin dosing (see Drug Update) and oxaliplatin bag size clarified		
GIGAVFFOX	Palliative Treatment of Metastatic or Locally Advanced Gastric, Gastroesophageal Junction or Esophageal Adenocarcinoma using Oxaliplatin, Fluorouracil and Leucovorin	Leucovorin dosing (see Drug Update) and oxaliplatin bag size clarified		
GIGAVFFOXT	Palliative Treatment of Metastatic or Locally Advanced Gastric, Gastroesophageal Junction or Esophageal Adenocarcinoma using Oxaliplatin, Fluorouracil, Leucovorin and Trastuzumab	Leucovorin dosing (see Drug Update) and oxaliplatin bag size clarified		

REVISED Protocols, PPPOs and Patient Handouts (revisions in respective columns)				
Code	Protocol Title	Protocol	РРРО	Handout
GIGAVTR	Continuation of Palliative Treatment of Metastatic or Inoperable, Locally Advanced Gastric or Gastroesophageal Junction Adenocarcinoma using Trastuzumab		Treatment updated (multiple cycles can be ordered)	
GIGFLODOC	Perioperative Treatment of Resectable Adenocarcinoma of the Stomach, Gastroesophageal Junction or Lower ½ Esophagus using Docetaxel, Oxaliplatin, Infusional Fluorouracil and Leucovorin	Treatment updated (filgrastim added); Leucovorin dosing (see Drug Update) and docetaxel and oxaliplatin bag sizes clarified	Premedication updated (filgrastim added)	Treatment Summary and Side Effects and Management updated (filgrastim added)
GIGFOLFIRI	Second-Line Palliative Combination Chemotherapy for Metastatic Gastric or Esophageal Adenocarcinoma using Irinotecan, Fluorouracil and Leucovorin	Leucovorin dosing clarified (see Drug Update)		
UGINETEV	Treatment of Advanced Neuroendocrine Tumours of Gastrointestinal Origin (Non- Functional) using Everolimus	Eligibility revised		
GIPAJFIROX	Adjuvant Chemotherapy for Resected Pancreatic Adenocarcinoma using Irinotecan, Oxaliplatin, Fluorouracil and Leucovorin	Leucovorin dosing (see Drug Update) and oxaliplatin bag size clarified	Pre-treatment tests revised (platelets)	
GIRAJFFOX	Adjuvant Combination Chemotherapy for Stage III Rectal Cancer using Oxaliplatin, Fluorouracil and Leucovorin	Leucovorin dosing (see Drug Update) and oxaliplatin bag size clarified		
GIREGO	Treatment of Advanced Hepatocellular Carcinoma using Regorafenib	Eligibility revised		
GO Gynecolo	gic			
GOOVBEVG	Treatment of Platinum-Resistant Ovarian Cancer with Bevacizumab and Gemcitabine	Treatment duration clarified		
GOOVBEVLD	Treatment of Platinum-Resistant Ovarian Cancer with Bevacizumab and Doxorubicin Pegylated Liposomal (CAELYX®)	Treatment duration clarified	Tests updated (AST replaced by ALT)	
GOOVBEVP	Treatment of Platinum-Resistant Ovarian Cancer with Bevacizumab and Paclitaxel	Treatment duration clarified	Tests updated (AST replaced by ALT)	
GOOVBEVV	Treatment of Platinum-Resistant Ovarian Cancer with Bevacizumab and Vinorelbine	Treatment duration clarified	Tests updated (AST replaced by ALT)	
GOOVCAG	Treatment of Advanced Ovarian Cancer in Patients Who Have Progressed or Recurred Following First-Line Platinum-Based Treatment using Carboplatin and Gemcitabine	Treatment duration clarified		

REVISED Protocols	DDDOs and Datient Handouts	(revisions in respective columns)
REVISED FIOLOCOIS,	rrrus and ratient nanduuts	(revisions in respective columns)

REVISED FIOLOCOIS, FFFOS and Fatient Handouts (revisions in respective columns)				
Code	Protocol Title	Protocol	РРРО	Handout
GOOVCARB	First- or Second-Line Therapy for Invasive Epithelial Ovarian Cancer using Single-Agent Carboplatin	Treatment duration clarified		
GOOVCATX	Primary Treatment of Visible Residual (Extreme Risk) Invasive Epithelial Ovarian, Fallopian Tube or Peritoneal Cancer using Carboplatin and Paclitaxel	Treatment duration clarified		
GOOVCYCPO	Palliative Therapy for Relapsed/Progressing Epithelial Ovarian, Primary Peritoneal or Fallopian Tube Carcinoma using Metronomic Low-Dose Oral Cyclophosphamide	Treatment duration clarified		
GOOVDOC	Treatment of Relapsed/Progressing Epithelial Ovarian, Primary Peritoneal or Fallopian Tube Carcinoma using Docetaxel	Treatment duration clarified		
GOOVETO	Treatment of Relapsed/Progressing Epithelial Ovarian, Primary Peritoneal or Fallopian Tube Carcinoma using Etoposide	Treatment duration clarified		
GOOVGEM	Treatment of Relapsed/Progressing Epithelial Ovarian, Primary Peritoneal or Fallopian Tube Carcinoma using Gemcitabine	Treatment duration clarified		
GOOVLDOX	Treatment of Epithelial Ovarian Cancer Relapsing after Primary Treatment using Doxorubicin Pegylated Liposomal (CAELYX®)	Treatment duration clarified		
GOOVTAX3	Treatment of Relapsed/Progressing Epithelial Ovarian, Primary Peritoneal or Fallopian Tube Carcinoma using Paclitaxel	Treatment duration clarified		
GOOVTOP	Treatment of Relapsed/Progressing Epithelial Ovarian, Primary Peritoneal or Fallopian Tube Carcinoma using Topotecan	Treatment duration clarified		
GOOVVIN	Treatment of Relapsed/Progressing Epithelial Ovarian, Primary Peritoneal or Fallopian Tube Carcinoma using Vinorelbine	Treatment duration clarified		
GU Genitourinary				
GUTAXGEM	Palliative Therapy for Germ Cell Cancers using Paclitaxel and Gemcitabine	Paclitaxel bag size clarified	Protocol Code corrected and paclitaxel bag size clarified	
HN Head and Neck				
HNOTLEN	Therapy for Locally Recurrent or Metastatic, RAI-Refractory Differentiated Thyroid Cancer using Lenvatinib	Tests corrected; Dose Modifications and Precautions clarified		

Code	Protocol Title	Protocol	РРРО	Handout
.U Lung				
LUAVCRIZ	Second-Line Treatment of ALK-Positive Advanced Non-Small Cell Lung Cancer with Crizotinib	Tests and Dose Modifications (hepatic) updated	Return Appointment tests updated	
LUAVCRIZF	First-Line Treatment of ALK-Positive Advanced Non-Small Cell Lung Cancer with Crizotinib	Tests and Dose Modifications (hepatic) updated	Return Appointment tests updated	
LUAVPMBM	Maintenance Therapy of Advanced Non-Small Cell Lung Cancer with Pembrolizumab	Eligibility clarified		
LUAVPMBM6	Maintenance Therapy of Advanced Non-Small Cell Lung Cancer with 6-Weekly Pembrolizumab	Eligibility clarified		
ULUAVPPPMB	First-Line Treatment of Advanced Non- Squamous Non-Small Cell Lung Cancer with Platinum, Pemetrexed and Pembrolizumab	Eligibility clarified		
LY Lymphom	a			
LYNIV	Treatment of Relapsed or Refractory Hodgkin Lymphoma using Nivolumab	Eligibility clarified		
LYNIV4	Treatment of Relapsed or Refractory Hodgkin Lymphoma using 4-Weekly Nivolumab	Eligibility clarified		
ULYPRA	Treatment of Relapsed or Refractory Peripheral T-Cell Lymphoma (PTCL) with Pralatrexate		Dose Modifications and telephone nursing assessment orders revised; "Start IV line" removed	
MY Myelom	a			
MYBORPRE	Treatment of Multiple Myeloma using Bortezomib, Dexamethasone with or without Cyclophosphamide as Induction Pre-Stem Cell Transplant	Prednisone dosing clarified		
MYBORREL	Treatment of Relapsed Multiple Myeloma using Bortezomib, Dexamethasone with or without Cyclophosphamide	Prednisone dosing clarified		
UMYCARDEX	Therapy of Multiple Myeloma using Carfilzomib and Dexamethasone with or without Cyclophosphamide		Cyclophosphamide dosing clarified	

Premedications revised (see Drug Update)

Treatment of Relapsed and Refractory Multiple Myeloma with Daratumumab in

Combination with Bortezomib and

Dexamethasone with or without

Cyclophosphamide

UMYDARBD

REVISED Protocols, PPPOs and Patient Handouts (revisions in respective columns)				
Code	Protocol Title	Protocol	РРРО	Handout
UMYDARLD	Treatment of Relapsed and Refractory Multiple Myeloma with Daratumumab in Combination with Lenalidomide and Dexamethasone	Premedications revised (see Drug Update)		
SC Supportiv	e Care			
SCNAUSEA	Guidelines for Prevention and Treatment of Chemotherapy-Induced Nausea and Vomiting in Adults	See Editor's Choice		
SM Skin and	Melanoma			
SMAVDT	Treatment of BRAF-Positive Unresectable or Metastatic Melanoma using Dabrafenib and Trametinib	AST deleted	AST deleted	
SMAVI	Treatment of Advanced C-Kit-Positive Melanoma using Imatinib	Institution name and Tests updated	Institution name and Tests updated	
SMAVNIV	Treatment of Unresectable or Metastatic Melanoma using Nivolumab	Contact Physician updated and Eligibility clarified		
SMAVNIV4	Treatment of Unresectable or Metastatic Melanoma using 4-Weekly Nivolumab	Contact Physician updated and Eligibility clarified		
SMAVPEM	Treatment of Unresectable or Metastatic Melanoma using Pembrolizumab	Contact Physician updated and Eligibility clarified		
SMAVPEM6	Treatment of Unresectable or Metastatic Melanoma using 6-Weekly Pembrolizumab	Contact Physician updated and Eligibility clarified		
SMAVTRA	Treatment of BRAF-Positive Unresectable or Metastatic Melanoma using Trametinib	AST deleted	AST deleted	
SMAVVC	Treatment of BRAF-Positive Unresectable or Metastatic Melanoma using Vemurafenib and Cobimetinib	AST deleted	Electrolytes clarified and AST deleted	
SMAVVEM	Treatment of BRAF-Positive Unresectable or Metastatic Melanoma using Vemurafenib	AST deleted	Electrolytes clarified and AST deleted	Institution name updated

The **CAP requirement** has been removed from the **Eligibility** section and the **U** removed from the **Protocol Code** in the following BC Cancer treatment protocols, provincial pre-printed orders and or patient handouts:

CODE	Protocol Title	
BR Breast		
⊎BRAJPAM	Adjuvant Therapy in Postmenopausal Women using Pamidronate	
⊎BRAJTTW	Adjuvant Therapy using Weekly Paclitaxel and Trastuzumab	
GI Gastrointe	istinal	
₩GIAJRALOX	Adjuvant Chemotherapy for Node-Positive Colon Cancer using Oxaliplatin and Raltitrexed in Patients Intolerant to Fluorouracil or Capecitabine	
₩GIDEBTACE	Transarterial Chemoembolization (TACE) of Hepatocellular Carcinoma using Drug-Eluting Bead (DEB) Loaded with Doxorubicin	
GO Gynecolo	gic	
⊎GOOVBEVG	Treatment of Platinum-Resistant Ovarian Cancer with Bevacizumab and Gemcitabine	
⊎GOOVBEVLD	Treatment of Platinum-Resistant Ovarian Cancer with Bevacizumab and Doxorubicin Pegylated Liposomal (CAELYX®)	
UGOOVBEVP	Treatment of Platinum-Resistant Ovarian Cancer with Bevacizumab and Paclitaxel	
UGOOVBEVV	Treatment of Platinum-Resistant Ovarian Cancer with Bevacizumab and Vinorelbine	
⊎GOOVCATB	Primary Treatment of Invasive Ovarian with High-Risk of Relapse using Bevacizumab, Carboplatin and Paclitaxel	
HN Head and	Neck	
₩HNLADCF	Locally Advanced Squamous Cell Carcinoma with Docetaxel, Cisplatin and Infusional Fluorouracil	
LY Lymphom	a	
UYBRENTUX	Treatment of Hodgkin Lymphoma and Anaplastic Large Cell Lymphoma with Brentuximab Vedotin	
ULYBV	Consolidation Therapy Post-ASCT for Hodgkin Lymphoma using Brentuximab Vedotin	
UUYIBRU	Treatment of Relapsed/Refractory Chronic Lymphocytic Leukemia using Ibrutinib	
	Treatment of Relapsed/Refractory Chronic Lymphocytic Leukemia using Idelalisib and Rituximab	
	Treatment of Cutaneous T-Cell Lymphoma (Mycosis Fungoides/Sézary Syndrome) with Bexarotene	
UYMIBRU	Treatment of Relapsed/Refractory Mantle-Cell Lymphoma using Ibrutinib	
ULYNIV	Treatment of Relapsed or Refractory Hodgkin Lymphoma using Nivolumab	
⊎LYNIV4	Treatment of Relapsed or Refractory Hodgkin Lymphoma using 4-Weekly Nivolumab	
ULYOBBEND	Treatment of Rituximab-Refractory Follicular Lymphoma with Obinutuzumab in Combination with Bendamustine	
	Treatment of Previously-Untreated Chronic Lymphocytic Leukemia with Obinutuzumab and Chlorambucil	
UYRITZ	Treatment of Lymphoma using Radioimmunotherapy: Rituximab-Priming for Ibritumomab ⁹⁰ Y (ZEVALIN®)	
	Treatment of Relapsed/Refractory Chronic Lymphocytic Leukemia using Venetoclax and Rituximab	
SM Skin and	Melanoma	
⊎SMAVDAB	Treatment of BRAF-Positive Unresectable or Metastatic Melanoma using Dabrafenib	
⊎SMAVDT	Treatment of BRAF-Positive Unresectable or Metastatic Melanoma using Dabrafenib and Trametinib	
⊎SMAVFIPI	First-Line Treatment of Unresectable or Metastatic Melanoma using Ipilimumab	
⊎SMAVI	Treatment of Advanced C-Kit-Positive Melanoma using Imatinib	
⊎SMAVIPI	Treatment of Unresectable or Metastatic Melanoma using Ipilimumab	
⊎SMAVNIV	Treatment of Unresectable or Metastatic Melanoma using Nivolumab	
⊎SMAVNIV4	Treatment of Unresectable or Metastatic Melanoma using 4-Weekly Nivolumab	
⊎ SMAVPEM	Treatment of Unresectable or Metastatic Melanoma using Pembrolizumab	
⊎SMAVPEM6	Treatment of Unresectable or Metastatic Melanoma using 6-Weekly Pembrolizumab	
⊎SMAVTRA	Treatment of BRAF-Positive Unresectable or Metastatic Melanoma using Trametinib	
₩SMAVVC	Treatment of BRAF-Positive Unresectable or Metastatic Melanoma using Vemurafenib and Cobimetinib	
⊎ SMAVVEM	Treatment of BRAF-Positive Unresectable or Metastatic Melanoma using Vemurafenib	
⊎ SMMCCAVE	Second-Line Treatment of Recurrent or Metastatic Merkel Cell Carcinoma using Avelumab	

Resources and Contact Information

Resource	Phone	Email / Toll Free / Fax	
Systemic Therapy Update: www.bccancer.bc.ca/health-professionals/clinical-resources/systemic-therapy/systemic-therapy-update			
Systemic Therapy Update Editor	604-877-6000 x 672649	bulletin@bccancer.bc.ca	
Oncology Drug Information Cancer Drug Manual Editor Pharmacy Oncology Certification Nurse Educators	604-877-6275 250-519-5500 x 693742 250-712-3900 x 686820 604-877-6000 x 672638	druginfo@bccancer.bc.ca nbadry@bccancer.bc.ca rxchemocert@bccancer.bc.ca nursinged@bccancer.bc.ca	
CAP – Compassionate Access Program	604-877-6277	cap bcca@bccancer.bc.ca fax 604-708-2026	
OSCAR – Online System for Cancer Drugs Adjudication and Reimbursement	888-355-0355	oscar@bccancer.bc.ca fax 604-708-2051	
Library/Cancer Information	604-675-8003	toll free 888-675-8001 x 8003 requests@bccancer.bc.ca	
Library Document Delivery	604-675-8002	requests@bccancer.bc.ca	
Pharmacy Professional Practice Professional Practice, Nursing Provincial Systemic Therapy Program	604-877-6000 x 672247 604-877-6000 x 672623 604-877-6000 x 672247	mlin@bccancer.bc.ca BCCancerPPNAdmin@ehcnet.phsa.ca mlin@bccancer.bc.ca	
BC Cancer – Abbotsford BC Cancer – Kelowna BC Cancer – Prince George BC Cancer – Surrey BC Cancer – Vancouver	604-851-4710 250-712-3900 250-645-7300 604-930-2098 604-877-6000	toll free 877-547-3777 toll free 888-563-7773 toll free 855-775-7300 toll free 800-523-2885 toll free 800-663-3333	
BC Cancer – Victoria	250-519-5500	toll free 800-670-3322	

Community Oncology Network (CON) sites: To update your contact information, please contact: bulletin@bccancer.bc.ca

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