



MEDICAL ONCOLOGY VALUE-BASED PATHWAYS

Evicore Oncology Value-Based Pathways

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EviCore healthcare Clinical Decision Support Tool Diagnostic Strategies: This tool addresses common symptoms and symptom complexes. Imaging requests for individuals with atypical symptoms or clinical presentations that are not specifically addressed will require physician review. Consultation with the referring physician, specialist, and/or individual's Primary Care Physician (PCP) may provide additional insight.

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Medical Oncology Value-Based Pathways

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EviCore's value-based pathways program was created to promote the use of evidence-based, high value, cancer treatment regimens. Pathway regimens are a subset of regimens recommended by the National Comprehensive Cancer Network[®] (NCCN) that deliver the best clinical outcomes for patients.

Evicore Oncology Value-Based Pathways

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Anal Cancer Pathways

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Localized Disease: Definitive Chemoradiation	Recurrent/Metastatic: First Line	Recurrent/Metastatic: Subsequent Therapy
• Mitomycin + fluorouracil (5FU)	• Carboplatin + paclitaxel	• Nivolumab
• Mitomycin + capecitabine		

Biliary Tract Cancers (Gallbladder, Extrahepatic, and Intrahepatic Cholangiocarcinoma) Pathways

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Unresectable/Metastatic: First Line	Unresectable/Metastatic: Second Line
• Durvalumab + gemcitabine + cisplatin	• FOLFOX (fluorouracil [5FU], leucovorin, and oxaliplatin)
• Gemcitabine + cisplatin	• FOLFIRI (fluorouracil [5FU], leucovorin, and irinotecan)
• Capecitabine	• Capecitabine
• Gemcitabine	• Gemcitabine

Bladder Cancer Pathways

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Neoadjuvant/ Adjuvant Therapy	Locally Advanced Unresectable or Metastatic: First Line, Cisplatin Eligible	Locally Advanced Unresectable or Metastatic: First Line, Cisplatin Ineligible	Locally Advanced/ Unresectable Metastatic: Second Line Post Chemotherapy and Enfortumab	Locally Advanced Unresectable Metastatic: Second Line Post Chemotherapy and Immunotherapy
• DDMVAC (dose-dense methotrexate, vinblastine, doxorubicin, and cisplatin)	• Pembrolizumab + enfortumab vedotin	• Pembrolizumab + enfortumab vedotin	• DDMVAC (dose-dense methotrexate, vinblastine, doxorubicin, and cisplatin)	• Enfortumab vedotin
• Gemcitabine + Cisplatin	• Nivolumab + gemcitabine + cisplatin	• Pembrolizumab	• Gemcitabine + Cisplatin	
	• Gemcitabine + cisplatin	• Gemcitabine + carboplatin	• Gemcitabine + carboplatin	
	• Avelumab maintenance (if no progression on platinum- based chemotherapy)		• Paclitaxel	
			• Docetaxel	
			• Gemcitabine	

Breast Cancer Pathways

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Neoadjuvant/Adjuvant

Neoadjuvant/Adjuvant: HER2 Negative	Triple Negative (HER2-, ER-, and PR-Negative) – 80kg or less patient
• TC (docetaxel and cyclophosphamide)	• Keynote 522 regimen with pembrolizumab weight-based dosing
• Dose Dense AC (doxorubicin and cyclophosphamide) followed by weekly paclitaxel	• NeoPact regimen with pembrolizumab weight-based dosing

Metastatic

HER2 Negative	HER2 Positive: First Line	HER2 Positive: Second Line or beyond	HER2 Positive: Third Line and beyond	HER2 Positive: Fourth Line or beyond
• Docetaxel	• Docetaxel + trastuzumab [*] + pertuzumab [§]	• Fam-trastuzumab deruxtecan	• Ado-trastuzumab emtansine	• Chemo + trastuzumab [*]
• Paclitaxel	• Paclitaxel + trastuzumab [*] + pertuzumab [§]		• Chemo + trastuzumab [*]	• Lapatinib + trastuzumab [*]
• Gemcitabine				
• Vinorelbine				
• Capecitabine				
• Doxorubicin				
• Liposomal doxorubicin				

* Use of preferred oncology medication products may be required for coverage. Refer to the health plan's coverage policy for additional information.

§ Pertuzumab, trastuzumab, and hyaluronidase-zzxf injection for subcutaneous use may be substituted for intravenous pertuzumab and intravenous trastuzumab.

Chronic Myeloid Leukemia Pathways

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First Line: Low Risk Score Disease

- Imatinib

Chronic Lymphocytic Leukemia/Small Lymphocytic Lymphoma Pathways

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First Line: WITHOUT del (17P)/TP53 Mutation	First Line: WITH del (17P)/TP53 Mutation
• Ibrutinib	• Ibrutinib
• Acalabrutinib	• Acalabrutinib
• Zanubrutinib	• Zanubrutinib
• Venetoclax + obinutuzumab	• Venetoclax + obinutuzumab
• Bendamustine + rituximab *	

* Use of preferred oncology medication products may be required for coverage. Refer to the health plan's coverage policy for additional information.

Colorectal Cancer Pathways

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Metastatic: First or Second Line Therapy for RAS/RAF Wild Type	Metastatic: Subsequent Therapy with a VEGF Inhibitor	Metastatic: dMMR/MSI-H or POLE/POLD1 Mutation
• Panitumumab-based therapy	• Bevacizumab* + FOLFIRI (fluorouracil [5FU], leucovorin, and irinotecan)	• Pembrolizumab
	• Bevacizumab* + irinotecan	

* Use of preferred oncology medication products may be required for coverage. Refer to the health plan's coverage policy for additional information.

Diffuse Large B-Cell Lymphoma Pathways

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Second Line Therapy, Relapsed Disease >12 Months: Intention to Proceed to Transplant	Second Line Therapy, Relapsed Disease >12 Months: No Intention to Proceed to Transplant
• ICE (ifosfamide, carboplatin and etoposide) +/- rituximab [*]	• GemOx (gemcitabine and oxaliplatin) +/- rituximab [*]
• GDP (gemcitabine, dexamethasone and cisplatin) +/- rituximab [*] or gemcitabine, dexamethasone and carboplatin) +/- rituximab [*]	• GDP (gemcitabine, dexamethasone and cisplatin) +/- rituximab [*] or gemcitabine, dexamethasone and carboplatin) +/- rituximab [*]
• DHAX (dexamethasone, cytarabine and oxaliplatin) +/- rituximab [*]	• Polatuzumab vedotin +/- bendamustine +/- rituximab [*]
	• Glofitamab + GemOx (gemcitabine and oxaliplatin)

* Use of preferred oncology medication products may be required for coverage. Refer to the health plan's coverage policy for additional information.

Endometrial Cancer Pathways

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Recurrent/Metastatic: First Line and Subsequent Therapy
• Carboplatin + paclitaxel
• Carboplatin + paclitaxel + trastuzumab [*] (only for HER2 positive serous carcinoma or carcinosarcoma)
• Cisplatin + doxorubicin
• Paclitaxel
• Doxorubicin
• Topotecan
• Cisplatin
• Carboplatin

^{*} Use of preferred oncology medication products may be required for coverage. Refer to the health plan's coverage policy for additional information.

Esophageal and Esophagogastric Junction Cancer Pathways

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Unresectable/Recurrent Metastatic: First Line, Adenocarcinoma, HER2 Negative, MSS	Unresectable/Recurrent Metastatic: First Line, Squamous Cell, MSS	Unresectable/Recurrent Metastatic: Subsequent Therapy, Adenocarcinoma or Squamous Cell
• FOLFOX (fluorouracil [5FU], leucovorin, and oxaliplatin)	• FOLFOX (fluorouracil [5FU], leucovorin, and oxaliplatin)	• Paclitaxel
• CapeOx (capecitabine and oxaliplatin)	• CapeOx (capecitabine and oxaliplatin)	• Docetaxel
• Fluorouracil [5FU] or capecitabine + cisplatin	• Fluorouracil [5FU] or capecitabine + cisplatin	• Irinotecan
• Nivolumab + FOLFOX (fluorouracil [5FU], leucovorin, and oxaliplatin) (For CPS ≥ 5 only)	• Nivolumab + (fluorouracil [5FU] or capecitabine) + cisplatin	• Pembrolizumab (For squamous cell, second line, CPS ≥ 10 and no prior CPI only)
• Nivolumab + CapeOx (capecitabine and oxaliplatin) (For CPS ≥ 5 only)	• Nivolumab + FOLFOX (fluorouracil [5FU], leucovorin, and oxaliplatin)	• Nivolumab (For squamous cell and no prior CPI only)
• Pembrolizumab + (fluorouracil [5FU] or capecitabine) + cisplatin (For CPS ≥ 10 only)	• Nivolumab + CapeOx (capecitabine and oxaliplatin)	
• Pembrolizumab + FOLFOX (fluorouracil [5FU], leucovorin, and oxaliplatin) (For CPS ≥ 10 only)	• Pembrolizumab + (fluorouracil [5FU] or capecitabine) + cisplatin (For CPS ≥ 10 only)	

Unresectable/Recurrent Metastatic: First Line, Adenocarcinoma, HER2 Negative, MSS	Unresectable/Recurrent Metastatic: First Line, Squamous Cell, MSS	Unresectable/Recurrent Metastatic: Subsequent Therapy, Adenocarcinoma or Squamous Cell
• Pembrolizumab + CapeOx (capecitabine and oxaliplatin) (<i>For CPS ≥ 10 only</i>)	• Pembrolizumab + FOLFOX (fluorouracil [5FU], leucovorin, and oxaliplatin) (<i>For CPS ≥ 10 only</i>)	
Tislelizumab + (fluorouracil [5FU] or capecitabine) + cisplatin (<i>For CPS ≥ 5 only</i>)	• Pembrolizumab + CapeOx (capecitabine and oxaliplatin) (<i>For CPS ≥ 10 only</i>)	
Tislelizumab + FOLFOX (fluorouracil [5FU], leucovorin, and oxaliplatin) (<i>For CPS ≥ 5 only</i>)	• Tislelizumab + (fluorouracil [5FU] or capecitabine) + cisplatin (<i>For CPS or TAP ≥ 10 only</i>)	
Tislelizumab + CapeOx (capecitabine and oxaliplatin) (<i>For CPS ≥ 5 only</i>)	• Tislelizumab + FOLFOX (fluorouracil [5FU], leucovorin, and oxaliplatin) (<i>For CPS or TAP ≥ 10 only</i>)	
	• Tislelizumab + CapeOx (capecitabine and oxaliplatin) (<i>For CPS or TAP ≥ 10 only</i>)	
	• Tislelizumab + cisplatin + paclitaxel (<i>For CPS or TAP ≥ 10 only</i>)	
	• Tislelizumab + oxaliplatin + paclitaxel (<i>For CPS or TAP ≥ 10 only</i>)	

Follicular B-Cell Lymphoma Pathways

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Grade 1-2 Disease: First Line
• BR (bendamustine and rituximab [*])
• R-CHOP (rituximab [*] , cyclophosphamide, doxorubicin, vincristine and prednisone)
• R-CVP (rituximab [*] , cyclophosphamide, vincristine and prednisone)

^{*} Use of preferred oncology medication products may be required for coverage. Refer to the health plan's coverage policy for additional information.

Head and Neck Cancer Pathways

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Advanced Disease: Definitive Chemoradiation
• High dose cisplatin
• Carboplatin + infusional fluorouracil (5FU)
• Weekly cisplatin
• Weekly carboplatin + paclitaxel

Hepatocellular Carcinoma Pathways

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Metastatic or Unresectable Disease: First line	Metastatic or Unresectable Disease: Subsequent Therapy
• Lenvatinib	• Regorafenib
• Atezolizumab + bevacizumab *	• Pembrolizumab
• Tremelimumab + durvalumab	
• Durvalumab	

* Use of preferred oncology medication products may be required for coverage. Refer to the health plan's coverage policy for additional information.

Hodgkin Lymphoma Pathways

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Stage III/IV Classical Hodgkin Lymphoma - Ages 18-60 years old

- Nivolumab-AVD (doxorubicin, vinblastine, and dacarbazine)

Kidney Cancer Pathways

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First line: Favorable Risk	Clear Cell, First Line: Intermediate or Poor Risk
• Lenvatinib + pembrolizumab	• Ipilimumab (1 mg/kg x 4 doses) + nivolumab
• Cabozantinib + nivolumab	
• Ipilimumab (1 mg/kg x 4 doses) + nivolumab	
• Sunitinib (<i>if not candidate for CPI</i>)	
• Pazopanib (<i>if not candidate for CPI</i>)	

Lung Cancer: Non-Small Cell Lung Cancer Pathways

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Metastatic, EGFR Exon 19 deletion or Exon 21 L858R Mutations

First Line	
• Osimertinib	
• Osimertinib + pemetrexed + (cisplatin or carboplatin) (<i>For nonsquamous only</i>)	

Nonsquamous, Metastatic

First Line, PD-L1 $\geq$ 50%, No Actionable Mutations	First line, PD-L1 < 50%, No Driver Mutations	Subsequent Therapy, No Actionable Mutations
• Pembrolizumab	• Pembrolizumab + pemetrexed + (carboplatin or cisplatin)	• Nivolumab (<i>only if no prior CPI</i>)
• Cemiplimab-rwlc	• Cemiplimab-rwlc + pemetrexed + (carboplatin or cisplatin)	• Pembrolizumab (<i>only if no prior CPI</i>)
• Atezolizumab	• Cemiplimab-rwlc + paclitaxel + (carboplatin or cisplatin)	• Atezolizumab (<i>only if no prior CPI</i>)
		• Pemetrexed
		• Gemcitabine
		• Docetaxel

Squamous, Metastatic

First Line, PD-L1 $\geq$ 50%, No Actionable Mutations	First line, PD-L1 < 50%, No Driver Mutations	Subsequent Therapy, No Actionable Mutations
• Pembrolizumab	• Pembrolizumab + paclitaxel + carboplatin	• Nivolumab (<i>only if no prior CPI</i>)

First Line, PD-L1 ≥ 50%, No Actionable Mutations	First line, PD-L1<50%, No Driver Mutations	Subsequent Therapy, No Actionable Mutations
• Cemiplimab-rwlc	• Cemiplimab-rwlc + paclitaxel + (carboplatin or cisplatin)	• Pembrolizumab (<i>only if no prior CPI</i>)
• Atezolizumab		• Atezolizumab (<i>only if no prior CPI</i>)
		• Gemcitabine
		• Docetaxel

Lung Cancer: Small Cell Lung Cancer Pathways

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Limited Stage Disease	Extensive Stage Disease: First Line	Extensive Stage Disease: Subsequent Therapy
• Cisplatin + etoposide +/- radiation	• Atezolizumab + carboplatin + etoposide	• Cisplatin + etoposide (<i>for prolonged disease-free interval</i>)
• Carboplatin + etoposide +/- radiation (<i>only if cisplatin ineligible or intolerant</i>)	• Carboplatin + etoposide	• Carboplatin + etoposide (<i>for prolonged disease-free interval</i>)
		• Topotecan

Myelodysplastic Syndrome (MDS) Pathways

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IPSS-R: Intermediate, High, Very High, Non-Transplant Candidate
• Azacitidine

Melanoma Pathways

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Stage III, Clinical Node Positive, Resectable Disease, Neoadjuvant/Adjuvant	Metastatic: First Line, BRAF Mutation Positive	Metastatic: First Line, BRAF Mutation Negative
• Nivolumab + ipilimumab followed by observation (if major pathological response)	• Vemurafenib + cobimetinib	• Pembrolizumab
	• Pembrolizumab	• Nivolumab
	• Nivolumab	• Ipilimumab + nivolumab
	• Ipilimumab + nivolumab	

Multiple Myeloma Pathways

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First Line: Transplant Eligible	First Line: Transplant Deferred or Not Indicated
• Bortezomib + lenalidomide + dexamethasone	• Bortezomib + lenalidomide + dexamethasone
• Daratumumab + lenalidomide + bortezomib + dexamethasone	• Lenalidomide + Dexamethasone
• Isatuximab + lenalidomide + bortezomib + dexamethasone	• Daratumumab + lenalidomide + dexamethasone
	• Daratumumab + lenalidomide + bortezomib + dexamethasone
	• Isatuximab + lenalidomide + dexamethasone
	• Isatuximab + lenalidomide + bortezomib + dexamethasone

Neuroendocrine Cancer Pathways

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Metastatic, Poorly-Differentiated, Large or Small Cell (Extrapulmonary)
• Cisplatin + etoposide
• Carboplatin + etoposide
• FOLFOX (fluorouracil [5FU], leucovorin, and oxaliplatin) (<i>following progression on platinum + etoposide</i>)
• FOLFIRI (fluorouracil [5FU], leucovorin, and irinotecan) (<i>following progression on platinum + etoposide</i>)

Ovarian Cancer Pathways

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Primary Therapy: Stage II-IV	Recurrent Disease, Platinum Sensitive
• Carboplatin + paclitaxel	• Carboplatin + gemcitabine
	• Cisplatin + gemcitabine
	• Carboplatin + gemcitabine

Pancreatic Cancer Pathways

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Adenocarcinoma: Neoadjuvant Therapy	Adenocarcinoma, Advanced/Metastatic: First Line Therapy
• Modified FOLFIRINOX (fluorouracil [5FU], leucovorin, irinotecan, and oxaliplatin)	• FOLFIRINOX or modified FOLFIRINOX
• Gemcitabine + cisplatin (<i>for BRCA1/2 or PALB2 mutations only</i>)	• Gemcitabine + albumin-bound paclitaxel

Prostate Cancer Pathways

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Metastatic: Adenocarcinoma, Castration Resistant
• Abiraterone
• Enzalutamide
• Docetaxel
• Cabazitaxel (post docetaxel)

Testicular Cancer Pathways

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First Line: Good Risk	First Line: Intermediate/Poor Risk
• BEP (bleomycin, etoposide, and cisplatin)	• BEP (bleomycin, etoposide, and cisplatin)

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References – Anal Cancer

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Localized Disease: Definitive Chemoradiation

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Recurrent/Metastatic: First Line

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Recurrent/Metastatic: Subsequent Therapy

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Unresectable/Metastatic: First Line

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References – Bladder Cancer

v2.0.2025

Neoadjuvant/Adjuvant Therapy

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Locally Advanced/Unresectable or Metastatic: First Line, Cisplatin Eligible

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References – Breast Cancer

v2.0.2025

Neoadjuvant/Adjuvant: HER2 Negative

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References – Chronic Lymphocytic Leukemia (CLL)

v2.0.2025

First Line, with or without del (17p)/TP53 Mutation

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References – Colorectal Cancer

v2.0.2025

Subsequent Therapy with a VEGF Inhibitor

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Metastatic Disease: dMMR/MSI-H or POLE/POLD1 Mutation

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Metastatic Disease: First Line or Second Line RAS/RAF Wild Type

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References – Diffuse Large B-Cell Lymphoma (DBCL)

v2.0.2025

Second Line Therapy, Relapsed Disease >12 months, No Intention to Proceed to Transplant or CART

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Second Line Therapy, Relapsed Disease >12 months, Intention to Proceed to Transplant

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References – Endometrial Cancer

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Recurrent/Metastatic: First Line and Subsequent Therapy

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References – Esophageal and Esophagogastric Junction (GEJ) Cancer

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Unresectable/Recurrent/Metastatic: First Line, Adenocarcinoma, HER2 Negative, MSS

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Grade 1-2 Disease: First Line

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References – Head and Neck Cancer

v2.0.2025

Advanced Disease: Definitive Chemoradiation

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References – Hepatocellular Carcinoma

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Metastatic or Unresectable Disease: First Line

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References – Hodgkin Lymphoma

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Stage III/IV Classical Hodgkin Lymphoma – Ages 18-60 years old

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References – Kidney Cancer

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Clear Cell, First Line: Favorable Risk

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Clear Cell, First Line: Intermediate/Poor Risk

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References – Lung Cancer: NSCLC

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Advanced/Metastatic: First Line, Squamous or Nonsquamous, PD-L1 $\geq$ 50%, No Actionable Mutations

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Metastatic: First Line, Squamous or Nonsquamous, PD-L1 <50%, No Driver Mutation

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Metastatic: Subsequent Therapy, Squamous or Nonsquamous, No Actionable Mutation

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Metastatic: First line, EGFR Exon 19 Deletion or Exon 21 L858R Mutations

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References – Lung Cancer: SCLC

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Extensive Stage Disease: First Line

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Extensive Stage Disease: Subsequent Therapy

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References – Myelodysplastic Syndrome (MDS)

v2.0.2025

IPSS-R: Intermediate, High, Very High, Non-Transplant Candidate

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References – Melanoma

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Stage III, Clinical Node Positive, Resectable Disease, Neoadjuvant/Adjuvant

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Metastatic: First Line Therapy BRAF Mutated

- Larkin J, Ascierto PA, Dreno B, et al. Combined vemurafenib and cobimetinib in BRAF-mutated melanoma. *N Engl J Med*. 2014;371(20):1867-1876. doi:10.1056/NEJMoa1408868.
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Metastatic: First Line Therapy, BRAF Mutation Negative

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References – Multiple Myeloma

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First Line: Transplant Eligible

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First Line: Transplant Deferred or Not Indicated

1. Durie BG, Hoering A, Abidi MH, et al. Bortezomib with lenalidomide and dexamethasone versus lenalidomide and dexamethasone alone in patients with newly diagnosed myeloma without intent for immediate autologous stem-cell transplant (SWOG S0777): a randomised, open-label, phase 3 trial. *Lancet*. 2017;389(10068):519-527. doi:10.1016/S0140-6736(16)31594-X.
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References – Neuroendocrine Cancer

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Metastatic, Poorly-Differentiated, Large or Small Cell (Extrapulmonary)

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References – Pancreatic Cancer

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Adenocarcinoma: Neoadjuvant

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Adenocarcinoma, Advanced/Metastatic: First Line

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References – Prostate Cancer

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Metastatic: Adenocarcinoma, Castration Resistant

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References – Testicular Cancer

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First Line: Intermediate/Poor Risk

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