

Systemic Therapy for SCLC: ASCO-OH (CCO) Guideline				
Clinical Question	Recommendation	Type	Evidence Quality	Strength
What is the optimal treatment regimen for adjuvant systemic therapy in patients with resected SCLC?	1.1. Adjuvant chemotherapy should be offered to patients with resected limited-stage SCLC who have adequate performance status.	EB	L	S
	1.2. Adjuvant chemotherapy should consist of 4 cycles of cisplatin or carboplatin plus etoposide.	IC	N/A	W
	1.3. Adjuvant chemotherapy should be initiated within 8 weeks from resection.	IC	N/A	W
What is the optimal systemic therapy for use with concurrent radiotherapy in patients with LS-SCLC?	2.1. Cisplatin and etoposide should be administered with concurrent radiotherapy in patients with LS-SCLC.	EB	H	S
	2.2. Carboplatin and etoposide may be offered as systemic therapy concurrent with radiation for patients with LS-SCLC and contraindications to the use of cisplatin.	EB	L	S
	2.3. Chemotherapy should be commenced as soon as possible in patients with LS-SCLC and not deferred until radiation therapy can be started.	IC	L	S
What is the optimal first-line systemic therapy for patients with ES-SCLC?	3.1. First-line systemic therapy with carboplatin or cisplatin plus etoposide plus immunotherapy (atezolizumab or durvalumab) followed by maintenance immunotherapy should be offered to patients with ES-SCLC if there are no contraindications to immunotherapy.	EB	H	S
What systemic therapy options are available for treating relapsed SCLC?	4.1. In patients with relapsed SCLC with a chemotherapy-free interval of less than 90 days, single-agent chemotherapy may be offered. Preferred agents are topotecan or lurbinectedin.	EB	M	S
	<i>Qualifying statement: Single-agent chemotherapy is preferred over multi-agent chemotherapy due to concerns regarding the balance of risks versus benefits.</i>			
	4.2. In patients with relapsed SCLC with a chemotherapy-free interval of at least 90 days, re-challenge with a platinum-based regimen or single-agent chemotherapy (preferred agents are topotecan or lurbinectedin) may be offered.	EB	M	S
	4.3. In patients with relapsed SCLC who had progression while on maintenance immunotherapy, there is no evidence to support continuation of immunotherapy.	IC	N/A	S

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	4.4. In an immunotherapy-naïve patient, second-line immunotherapy alone is not recommended outside of the clinical trial setting. Participation in clinical trials to better identify predictive biomarkers is encouraged.	EB	M	S
What is the best management approach for treatment naïve patients who are older or who have poor PS?	5.1. Older patients with LS-SCLC and ECOG PS 0-1 may be offered standard treatment with concurrent chemoradiotherapy with curative intent.	EB	M	S
	5.2. Patients with LS-SCLC and ECOG PS 2 due to SCLC may be offered standard treatment with concurrent chemoradiotherapy with curative intent.	EB	L	W
	5.3. Patients with LS-SCLC and ECOG PS 3-4 due to SCLC may be offered initial chemotherapy followed by sequential radiotherapy if there is improvement in PS.	IC	L	W
	5.4. Older patients with ES-SCLC and ECOG PS 0-1 may be offered standard treatment with carboplatin and etoposide plus immunotherapy (atezolizumab or durvalumab) followed by maintenance immunotherapy.	EB	M	S
	5.5. Patients with ES-SCLC and ECOG PS 2 may be offered carboplatin and etoposide plus immunotherapy.	IC	L	W
	5.6. Patients with ES-SCLC and ECOG PS 3-4 due to SCLC may be offered chemotherapy.	IC	L	W
What is optimal systemic therapy for patients with NSCLC harboring an EGFR mutation that has transformed to SCLC?	6.1. Patients with NSCLC harboring an <i>EGFR</i> mutation that has transformed to SCLC should be managed with carboplatin or cisplatin plus etoposide.	IC	L	W
	<i>Qualifying statement: There is insufficient evidence to support the use of immunotherapy in this setting. Clinical trial enrollment should be offered whenever possible.</i>			
	6.2. EGFR inhibitor may be continued with chemotherapy in patients with NSCLC harboring an <i>EGFR</i> mutation that has transformed to SCLC.	IC	L	W
What is the role of biomarkers, including molecular profiling in guiding therapy for patients with SCLC?	7.1. There is no evidence to support the use of molecular profiling and biomarker analysis to guide standard treatment in patients with <i>de novo</i> SCLC.	EB	L	W
Which myeloid supportive agents may be considered for use in patients with SCLC?	8.1. Trilaciclib or G-CSF may be offered as a myeloid supportive agent for patients with untreated or previously treated ES-SCLC who are undergoing treatment with chemotherapy or chemoimmunotherapy.	EB	M	W
	8.2. G-CSF may be offered in patients with LS-SCLC who are undergoing chemoradiotherapy.	EB	M	W

Abbreviations. ASCO, American Society of Clinical Oncology; CCO, Cancer Care Ontario; EB, evidence based; ECOG, Eastern Cooperative Oncology Group; EGFR, epidermal growth factor receptor; ES-SCLC, extensive-stage small-cell lung cancer; G-CSF, granulocyte colony stimulating factor; H, high; IC, informal consensus; L, low; LS-SCLC, limited-stage small-cell lung cancer; M, moderate; N/A, not applicable; NSCLC, non-small-cell lung cancer; OH, Ontario Health; PS, performance status; S, strong; SCLC, small-cell lung cancer; W, weak