

Radioactive iodine and other thyroid cancer treatments, and their side effects: frequently asked questions

How are patients prepared for radioactive iodine (RAI)?

Table 1 summarises the preparation of patients who will be treated with RAI (131]).

Increase thyroid stimulating hormone (TSH) levels to >30 mIU/L	 Increasing TSH levels ensures that RAI is maximally taken up by thyroid cells. This can be achieved by: Injecting recombinant human TSH 24 and 48 hours before RAI, while the patient receives levothyroxine¹ Withdrawing thyroid hormone²
Reduce iodine stores in the body	 This can be achieved by: Following a low iodine diet for 1–2 weeks before RAI²; (this advice can differ between centres) Not performing imaging using RAI contrast medium during 8 weeks before RAI² Discontinuing medications that are high in iodine²
Provide patients with support and advice	 Education, support and advice could include: Explaining the rationale for the above steps and the potential side effects of RAI^{2.3} Advising what to bring into the isolation room Advising about pregnancy and breast feeding (see below)² Highlighting the importance of avoiding close contact with others for a time after receiving RAI² and other practicalities after discharge

Table 1: Preparation of patients for RAI

Which patients are unsuitable for RAI?

RAI scans are contraindicated during pregnancy and breastfeeding. Healthcare professionals (HCPs) must exclude pregnancy before radioiodine remnant ablation (RRA) or RAI in women of reproductive potential. All patients (regardless of gender) should not attempt conception for at least 6 months after RRA or 4 months after RAI. Women should stop breastfeeding at least 8 weeks before RRA or RAI and not resume until after a future pregnancy.²





How should persistently increases in thyroglobulin (Tg) or Tg antibody (TgAb) levels in differentiated thyroid cancer be managed?

Rising Tg or TgAb levels may indicate the cancer's progression² and, therefore, indicate that further investigations, such as imaging, are needed. These investigations will indicate if further therapy, such as local excision or localised external beam radiation therapy, is required.¹² RAI will typically continue until the disease becomes RAI-refractory, but the dose may need to be increased.¹Expert opinion suggests continuing TSH suppression while closely monitoring Tg levels if the thyroid cancer becomes RAI-refractory and the patient is asymptomatic.

What are common post-operative complications?

Post-operative complications are generally rare. A study of 27,912 people who underwent thyroidectomy for differentiated or medullary thyroid cancer reported that 6.5% developed general surgical complications within 30 days of the procedure. During the year after the procedure, 12.3% of patients experienced complications specific to thyroidectomy.⁴

A review of surgery for thyroid cancer noted that:

- The most serious complications of thyroidectomy are injury to the recurrent laryngeal nerve (RLN) and hypocalcaemia.⁵
- Up to 5% of people show temporary nerve palsy following thyroidectomy, while less than 2–3% experience permanent RLN injury and vocal cord paralysis.⁵
- Bilateral RLN damage occurs after about 1 in 1000 thyroidectomies but requires permanent tracheotomy.⁵

Less common important complications include:

- Neck haematoma (<1% of cases).⁵
- Seroma.⁵
- Injury to the external branch of the superior laryngeal nerve (SLN). About a third of patients experience vocal symptoms following thyroidectomy, even without obvious nerve damage. Half of these still have symptoms 3 months after thyroidectomy, which may suggest SLN injury, especially among patients with, for example, reduced tone, vocal fatigue and problems projecting their voice.⁵
- Wound infections (e.g. cellulitis), while rare (0.1%-2%), are most common in immunocompromised patients and those with diabetes.⁵⁻⁷

What is the post-operative advice to patients?

Complications following thyroidectomy performed by experienced surgeons are generally uncommon and can be corrected.⁶ Most scars are barely visible six months to a year after thyroidectomy. Some patients, particularly those with red hair and those from Africa or the West Indies, can develop a hypertrophic (keloid) scar. Steroid tape and injections, and laser treatment may improve the appearance of keloid scare.⁸

Patients who undergo thyroidectomy should inform their HCP immediately if they experience muscle cramping or tingling in their hands, fingers or face, especially the lips.^{8,9} Adequate calcium concentrations are essential for normal nerve and muscle functioning.¹⁰ These symptoms may indicate low calcium levels and patients may need supplements.^{8,9} HCPs should also stress the importance of:

- Attending the follow up appointments and visits for thyroid function monitoring.⁸
- Reporting any symptoms that may suggest hypothyroidism.⁸
- Seeking urgent medical attention if the neck swelling seems excessive or they experience problems breathing.⁹





What are the main side-effects of tyrosine kinase inhibitors (TKI) and how can they be managed?

Table 2 summarises main side-effects associated with TKIs and their management. If the specific measures in table 2 do not alleviate or control the side effects, the HCP can consider adjusting the dose and duration of the TKI.^{11,12} The next question considers skin problems that may be associated with TKIs.

Hypertension	Ensure blood pressure is normal before starting TKI treatment			
	Aim to maintain a blood pressure of 140/90 mmHg or less, using anti-hypertensives if blood pressure exceeds this threshold			
Zz Z Fatigue	Encourage a nutritious diet and optimal hydration, with regular mild-moderate exercise			
	Recommend energy management strategies, such as rest periods during the day			
	Support the patient to manage other TKI side effects			
	Investigate other potential factors that may contribute to fatigue			
	Consider TKI administration in the evening, which may reduce daytime fatigue			
Diarrhoea	Regularly assess diarrhoea frequency and monitor electrolytes			
	Encourage adequate fluid intake			
	Avoid foods and drinks that could contribute to diarrhoea			
	Prescribe anti-diarrhoea medication			
Decreased appetite and weight loss	Encourage a nutritious diet			
	Monitor appetite and weight loss at each assessment			
	Treat any underlying nausea			

Table 2: Prophylactic measures and side effect Management for patients taking TKIs^{11,12}





What is the management of skin problems associated with TKIs?

TKIs can cause several dermatological adverse events including:

- Palmar-plantar erythrodysaesthesia (hand-foot) syndrome (PPES).¹³
- Rash (e.g. acneiform).¹³
- Erythema.13
- Pruritus.13
- Paronychia.¹³
- Telangiectasia.13
- Alopecia and changes in hair growth or pigmentation. ¹³
- Skin discoloration and dryness (xerosis). ¹³

Many dermatological adverse events can be treated or prevented (tables 3 and 4). HCPs should advise patients that acneiform rash and PPES generally develop within the first 8 weeks of treatment. Sun protection includes using a non-alcohol-based sunblock (SPF≥30) applied at least every two hours or every hour if sweating or swimming, protective clothing, including a hat, and avoiding direct sunlight. ¹³



Limit time in baths and showers; use cool or warm (not hot) water

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Use mild soaps and low pH cleansers and moisturisers; avoid cleansers containing alcohol



Apply moisturisers twice daily; apply after bathing while skin is damp



Reduce skin friction; pat rather than rub with a towel



Maintain humidity (reduce temperature in the home; use a humidifier)



Wear gloves and socks when exercising

Keep fingernails short



Wear hat and gloves in cold weather



Use fragrance-free and vitamin-free lip balm



Use moisturisers containing salicylic acid or ammonium lactate for very dry, scaly skin or xerosis of the palms or soles





	Grade 0	Grade 1	Grade 2	Grade 3*	Grade 4
Reaction	No adverse events	Mild; asymptomatic or mild symptoms; clinical or diagnostic observations only	Moderate; minimal, local or noninvasive intervention needed; limits age-appropriate instrumental activities of daily living (ADL) [§]	Severe or medically significant but not immediately life-threatening; hospitalisation or prolongation of hospitalisation; disabling; limiting self-care ADL [¶]	Life-threatening consequences; urgent intervention needed
Photosensitivity		Topical steroids	Topical steroids	Topical steroids	Oral antihistamines
	Sun protection	OTC moisturisers ⁺	Oral antihistamines	Oral antihistamines	Oral steroids
				Oral steroids	
PPES	Urea cream	Urea cream	Urea cream	Clobetasol cream	
	Ammonia lactate cream	Clobetasol cream	Clobetasol cream	Lidocaine (topical or patch)	
			Lidocaine (topical or patch)	Analgesia (NSAIDs; GABA agonists; narcotics)	
	Heavy moisturiser		Analgesia (NSAIDs; GABA agonists; narcotics)		
Acneiform rash	Oral doxycycline or oral minocycline [‡]	Hydrocortisone cream	Hydrocortisone cream	Hydrocortisone cream	Not stated
	Low-potency topical steroid [‡]			Oral doxycycline or oral minocycline	
	Sunscreen (SPF ≥30) [‡]		Oral doxycycline or oral		
	Moisturiser cream [‡]	Clindamycin gel	minocycline	Oral prednisolone	
	Skin care advice				

* Or intolerable grade 2; ⁺ containing aloe vera or pramoxine; ⁺ Prophylactic; ⁵ preparing meals, shopping for groceries or clothes, using the telephone, managing money, etc; ⁴ bathing, dressing and undressing, feeding self, using the toilet, taking medications, and not bedridden. Please see reference for specific recommendations (e.g. dose), monitoring, discontinuation and referral¹³ Paper uses National Cancer Institute Common Terminology Criteria for Adverse Events v 4.0.¹⁴ NSAID: Non-steroidal anti-inflammatory drug; OTC: Over-the-counter; PPES: palmar-plantar erythrodysaesthesia (hand-foot) syndrome

Table 4: Management of dermatological adverse events in patients taking TKI^{13,14}





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