

Myeloma: diagnosis and management

NICE guideline

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Your responsibility

The recommendations in this guideline represent the view of NICE, arrived at after careful consideration of the evidence available. When exercising their judgement, professionals and practitioners are expected to take this guideline fully into account, alongside the individual needs, preferences and values of their patients or the people using their service. It is not mandatory to apply the recommendations, and the guideline does not override the responsibility to make decisions appropriate to the circumstances of the individual, in consultation with them and their families and carers or guardian.

All problems (adverse events) related to a medicine or medical device used for treatment or in a procedure should be reported to the Medicines and Healthcare products Regulatory Agency using the [Yellow Card Scheme](#).

Local commissioners and providers of healthcare have a responsibility to enable the guideline to be applied when individual professionals and people using services wish to use it. They should do so in the context of local and national priorities for funding and developing services, and in light of their duties to have due regard to the need to eliminate unlawful discrimination, to advance equality of opportunity and to reduce health inequalities. Nothing in this guideline should be interpreted in a way that would be inconsistent with complying with those duties.

Commissioners and providers have a responsibility to promote an environmentally sustainable health and care system and should [assess and reduce the environmental impact of implementing NICE recommendations](#) wherever possible.

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This guideline is the basis of QS150.

Overview

This guideline covers the diagnosing and managing of myeloma (including smouldering myeloma and primary plasma cell leukaemia) in people aged 16 and over. It aims to improve care for people with myeloma by promoting the most effective tests and treatments for myeloma and its complications.

Who is it for?

- Healthcare professionals responsible for diagnosing and managing myeloma
- Hospitals that provide myeloma management
- Commissioners of specialised services
- People with myeloma, their families and carers

Recommendations

People have the right to be involved in discussions and make informed decisions about their care, as described in [NICE's information on making decisions about your care](#).

[Making decisions using NICE guidelines](#) explains how we use words to show the strength (or certainty) of our recommendations, and has information about prescribing medicines (including off-label use), professional guidelines, standards and laws (including on consent and mental capacity), and safeguarding.

Healthcare professionals should also follow our guidance on:

- [Haematological cancers: improving outcomes](#)
- [Improving outcomes in children and young people with cancer](#)
- [Patient experience in adult NHS services](#)
- [Shared decision making](#)

1.1 Communication and support

1.1.1 Consider providing the following information to people with myeloma:

- the symptoms of myeloma and treatment-related side effects (including steroid-related side effects, infection and neuropathy)
- lifestyle measures to optimise bone health and renal function
- how to identify and report new symptoms (especially pain and spinal cord compression).

1.2 Laboratory investigations

Laboratory investigations for people with suspected myeloma

For guidance on the setup of laboratory diagnostic services, see the [section on integrated diagnostic reporting in NICE's guideline on haematological cancers](#).

- 1.2.1 Use serum protein electrophoresis and serum-free light-chain assay to confirm the presence of a paraprotein indicating possible myeloma or monoclonal gammopathy of undetermined significance (MGUS).
- 1.2.2 If serum protein electrophoresis is abnormal, use serum immunofixation to confirm the presence of a paraprotein indicating possible myeloma or MGUS.
- 1.2.3 Do not use serum protein electrophoresis, serum immunofixation, serum-free light-chain assay or urine electrophoresis (urine Bence–Jones protein assessment) alone to exclude a diagnosis of myeloma.
- 1.2.4 When doing a bone marrow aspirate and trephine biopsy to confirm a diagnosis of myeloma, use morphology to determine plasma cell percentage and flow cytometry to determine plasma cell phenotype.

Laboratory investigations to provide prognostic information

- 1.2.5 Use the same sample for all diagnostic and prognostic tests on bone marrow, so people only need to have one bone marrow aspirate and trephine biopsy.
- 1.2.6 When doing a bone marrow aspirate and trephine biopsy to provide prognostic information:
 - Perform fluorescence in-situ hybridisation (FISH) on CD138-selected bone marrow plasma cells to identify the adverse risk abnormalities t(4;14), t(14;16), 1q gain, del(1p) and del(17p) (TP53 deletion). Use these abnormalities alongside International Staging System (ISS) scores to identify people with high-risk myeloma.

- Consider performing FISH on CD138-selected bone marrow plasma cells to identify the adverse risk abnormality t(14;20), and the standard risk abnormalities t(11;14) and hyperdiploidy.
 - Consider performing immunophenotyping of bone marrow to identify plasma cell phenotype, and to inform subsequent monitoring.
 - Consider performing immunohistochemistry (including Ki-67 staining and p53 expression) on the trephine biopsy to identify plasma cell phenotype and give an indication of cell proliferation, to provide further prognostic information.
- 1.2.7 Perform serum-free light-chain assay and use serum-free light-chain ratio to assess prognosis.

1.3 Imaging investigations

Imaging for people with suspected myeloma

- 1.3.1 Offer imaging to all people with a plasma cell disorder suspected to be myeloma.
- 1.3.2 Consider whole-body MRI as first-line imaging.
- 1.3.3 Consider whole-body low-dose CT as first-line imaging if whole-body MRI is unsuitable or the person declines it.
- 1.3.4 Consider skeletal survey as first-line imaging only if whole-body MRI and whole-body low-dose CT are unsuitable or the person declines them.
- 1.3.5 Do not use isotope bone scans to identify myeloma-related bone disease in people with a plasma cell disorder suspected to be myeloma.

Imaging for people with newly diagnosed myeloma

For guidance on imaging for people with suspected spinal cord compression, see the [section on imaging investigations in NICE's guideline on spinal metastases and metastatic spinal cord compression](#).

1.3.6 For people with newly diagnosed myeloma or smouldering myeloma who have not had whole-body imaging with 1 of the following, consider whole-body imaging to assess for myeloma-related bone disease and extra-medullary plasmacytomas with one of:

- MRI
- CT
- fluorodeoxyglucose positron emission tomography CT (FDG PET-CT).

1.3.7 Consider baseline whole-body imaging with MRI or FDG PET-CT for people who have non-secretory myeloma or suspected or confirmed soft tissue plasmacytomas and have not already had either of these tests.

1.4 Service organisation

For guidance on the facilities needed to provide intensive inpatient chemotherapy and transplants for people with myeloma, and the structure and function of multidisciplinary teams, see [NICE's guideline on improving outcomes in haematological cancers](#).

For guidance on service organisation for young people, see [NICE's cancer service guideline on improving outcomes in children and young people with cancer](#).

1.4.1 Each hospital treating people with myeloma who are not receiving intensive inpatient chemotherapy or a transplant should provide local access to:

- a multidisciplinary team specialising in myeloma
- supportive and palliative care, supported by:
 - psychological support services
 - a 24-hour acute oncology and/or haematology helpline
 - physiotherapy
 - occupational therapy

- dietetics
- medical social services
- critical care
- clinical trials via the MDT specialising in myeloma
- dental services.

1.4.2 Each hospital treating myeloma should provide regional access through its network to:

- facilities for intensive inpatient chemotherapy or transplantation
- renal support
- spinal disease management
- specialised pain management
- therapeutic apheresis
- radiotherapy
- restorative dentistry and oral surgery
- clinical trials, in particular early phase trials.

1.5 Managing newly diagnosed myeloma

First-line treatment

NICE has a suite of technology appraisal guidance on myeloma either published or in development. These published technology appraisals cover NICE's position in relation to primary disease treatment, salvage therapy for relapsed myeloma and consolidation/maintenance therapy after primary management. The recommendations in this guideline complement the existing technology appraisals, giving further guidance in addition to the technology appraisals where myeloma-related subgroups are not included.

- 1.5.1 Bortezomib is recommended as an option within its marketing authorisation, that is, in combination with dexamethasone, or with dexamethasone and thalidomide, for the induction treatment of adults with previously untreated multiple myeloma, who are eligible for high-dose chemotherapy with haematopoietic stem cell transplantation. [This recommendation is from [NICE technology appraisal guidance on bortezomib for induction therapy in multiple myeloma before high-dose chemotherapy and autologous stem cell transplantation.](#)]
- 1.5.2 Thalidomide in combination with an alkylating agent and a corticosteroid is recommended as an option for the first-line treatment of multiple myeloma in people for whom high-dose chemotherapy with stem cell transplantation is considered inappropriate. [This recommendation is from [NICE technology appraisal guidance on bortezomib and thalidomide for the first-line treatment of multiple myeloma.](#)]
- 1.5.3 Bortezomib in combination with an alkylating agent and a corticosteroid is recommended as an option for the first-line treatment of multiple myeloma if:
- high-dose chemotherapy with stem cell transplantation is considered inappropriate and
 - the person is unable to tolerate or has contraindications to thalidomide. [This recommendation is from [NICE technology appraisal guidance on bortezomib and thalidomide for the first-line treatment of multiple myeloma.](#)]

First autologous stem cell transplantation

- 1.5.4 Consider using frailty and performance status measures that include comorbidities to assess the suitability of first autologous stem cell transplant for people with myeloma.
- 1.5.5 Do not use age or the level of renal impairment alone to assess the suitability of first autologous stem cell transplant for people with myeloma.

Allogeneic stem cell transplantation

- 1.5.6 Take into account that allogeneic stem cell transplantation is only suitable for a small number of people with myeloma.
- 1.5.7 When assessing suitability of an allogeneic stem cell transplant for myeloma, take into account:
- whether the person has chemosensitive disease
 - how many previous lines of treatment they have had
 - whether a fully human leukocyte antigen (HLA) matched donor is available
 - how graft-versus-host disease (GvHD) and other complications may get worse with age
 - the risk of higher transplant-related mortality and morbidity, versus the potential for long-term disease-free survival
 - how outcomes could be improved with other, newer treatments
 - the person's understanding of the procedure and its risks and benefits.
- 1.5.8 Consider allogeneic stem cell transplantation as part of a clinical trial if one is available.

Primary plasma cell leukaemia

- 1.5.9 Consider bortezomib-based, lenalidomide-based, or both, combination induction chemotherapy for people with primary plasma cell leukaemia.
- 1.5.10 Consider high-dose melphalan-based autologous stem cell transplantation for people with primary plasma cell leukaemia if it is suitable.

1.6 Managing acute renal disease caused by myeloma

- 1.6.1 Consider immediately starting a bortezomib- and dexamethasone-based combination regimen for people with untreated, newly diagnosed, myeloma-induced acute renal disease.
- 1.6.2 If a bortezomib-based combination regimen is unsuitable for people with untreated, newly diagnosed, myeloma-induced acute renal disease, consider immediately starting a thalidomide- and dexamethasone-based combination regimen.

In February 2016, this was an off-label use. See [NICE's information on prescribing medicines](#).

- 1.6.3 Do not do plasma exchange for myeloma-induced acute renal disease.

1.7 Preventing and managing bone disease

Preventing bone disease

- 1.7.1 To prevent bone disease, offer people with myeloma:
- zoledronic acid **or**
 - disodium pamidronate, if zoledronic acid is contraindicated or not tolerated **or**
 - sodium clodronate, if zoledronic acid and disodium pamidronate are contraindicated, not tolerated or not suitable.
- 1.7.2 Consider immediately referring people with myeloma for dental assessment and treatment before starting zoledronic acid or disodium pamidronate.
- 1.7.3 For people who need urgent myeloma treatment, consider referring for dental assessment and treatment as soon as possible after they start treatment.

Managing non-spinal bone disease

- 1.7.4 Offer people with myeloma and non-spinal bone disease who have not already started bisphosphonates:
- zoledronic acid **or**
 - disodium pamidronate, if zoledronic acid is contraindicated or not tolerated **or**
 - sodium clodronate, if zoledronic acid and disodium pamidronate are contraindicated, not tolerated or not suitable.
- 1.7.5 Assess the risk of fracture (in line with the [NICE guideline on assessing the risk of fragility fractures in osteoporosis](#)) in people with myeloma and non-spinal bone disease.
- 1.7.6 Consider surgical stabilisation followed by radiotherapy for non-spinal bones that have fractured or are at high risk of fractures.
- 1.7.7 Consider radiotherapy for non-spinal bones that have fractured or are at high risk of fracture if surgical intervention is unsuitable or not immediately needed.
- 1.7.8 Consider radiotherapy for people with myeloma and non-spinal bone disease who need additional pain relief if:
- chemotherapy and initial pain management has not led to prompt improvement in pain control
 - chemotherapy is unsuitable and current pain medication is not working.
- 1.7.9 Consider re-treatment with radiotherapy if pain recurs or if there is regrowth of a previously treated lesion.
- 1.7.10 Consider seeking advice from or referral to specialists in palliative care or pain medicine for people with complex non-spinal bone disease.

Managing spinal bone disease

- 1.7.11 For guidance on treating metastatic spinal cord compression, see the [NICE guideline on metastatic spinal cord compression](#).
- 1.7.12 Offer all people with myeloma and spinal bone disease:
- bisphosphonates as follows, if not already started:
 - zoledronic acid **or**
 - disodium pamidronate, if zoledronic acid is contraindicated or not tolerated **or**
 - sodium clodronate, if zoledronic acid and disodium pamidronate are contraindicated, not tolerated or unsuitable
 - systemic pain control, when relevant, using the [NICE guidelines on neuropathic pain](#) and [opioids in palliative care](#).
- 1.7.13 Consider the following, in addition to other treatments, for all people with myeloma and spinal bone disease:
- interventional pain management
 - bracing.
- 1.7.14 For people with radiological evidence of myeloma-related spinal instability, consider immediate intervention with:
- spinal surgery, with or without radiotherapy
 - cement augmentation, with or without radiotherapy
 - radiotherapy alone, if spinal intervention is unsuitable or not currently needed.
- 1.7.15 In people with radiological evidence of myeloma-related spinal bone disease without instability, consider:
- cement augmentation, with or without radiotherapy

- radiotherapy alone.

1.8 Preventing and managing complications

Preventing infection

- 1.8.1 Offer people with myeloma the seasonal influenza vaccination.
- 1.8.2 Consider extending the pneumococcal vaccination to people with myeloma who are under 65.
- 1.8.3 Consider intravenous immunoglobulin replacement therapy for people who have hypogammaglobulinaemia and recurrent infections.
- 1.8.4 Consider continuing aciclovir or equivalent antiviral prophylaxis after treatment with bortezomib or other proteasome inhibitors ends.

In February 2016, this was an off-label use. See [NICE's information on prescribing medicines](#).

- 1.8.5 Consider aciclovir or equivalent antiviral prophylaxis for people who are taking both immunomodulatory drugs and high-dose steroids.

In February 2016, this was an off-label use. See [NICE's information on prescribing medicines](#).

- 1.8.6 Consider testing for hepatitis B, hepatitis C and HIV before starting myeloma treatment.

Managing peripheral neuropathy

- 1.8.7 For guidance on the pharmacological management of neuropathic pain see the [NICE guideline on neuropathic pain in adults](#).

- 1.8.8 Explain the symptoms of neuropathy to people with myeloma, and encourage them to tell their clinical team about any new, different or worsening neuropathic symptoms immediately.
- 1.8.9 If people who are receiving bortezomib develop neuropathic symptoms, consider immediately doing 1 or more of the following:
- switching to subcutaneous injections
 - reducing to weekly doses
 - reducing the dose.
- 1.8.10 Consider reducing the dose if people are taking a drug other than bortezomib and develop neuropathic symptoms.
- 1.8.11 Temporarily stop neuropathy-inducing myeloma treatments if people develop either of the following:
- grade 2 neuropathy with pain
 - grade 3 or 4 neuropathy.
- 1.8.12 If neuropathy does not improve despite stopping myeloma treatment and further treatment is needed, consider switching to myeloma treatments less likely to induce neuropathy.

Preventing thrombosis

Follow the recommendations in [NICE's guideline on venous thromboembolism in over 16s](#).

Managing fatigue

- 1.8.13 If other treatable causes of anaemia have been excluded, consider erythropoietin analogues (adjusted to maintain a steady state of haemoglobin at 110 to 120 g/litre) to improve fatigue in people with myeloma who have symptomatic anaemia.

1.9 Monitoring

- 1.9.1 Monitor smouldering myeloma every 3 months for the first 5 years, and then decide the frequency of further monitoring based on the long-term stability of the disease.
- 1.9.2 Monitor myeloma at least every 3 months in people who have completed treatment for the condition and have recovered. Take into account any risk factors for progression, such as:
- high-risk fluorescence in-situ hybridisation (FISH)
 - impaired renal function
 - disease presentation.
- 1.9.3 Monitoring for myeloma and smouldering myeloma should include:
- assessment of symptoms related to myeloma and myeloma treatment and
 - the following laboratory tests:
 - full blood count
 - renal function
 - bone profile
 - serum immunoglobulins and serum protein electrophoresis
 - serum-free light-chain assay, if appropriate.
- 1.9.4 Do not offer people with myeloma or smouldering myeloma routine skeletal surveys for disease monitoring.
- 1.9.5 Consider symptom-directed imaging for people with myeloma or smouldering myeloma if any new bone symptoms develop.
- 1.9.6 For people with myeloma and serological relapse or disease progression, consider 1 of the following (taking into consideration previous imaging tests):

- whole-body MRI
- spinal MRI
- fluorodeoxyglucose positron emission tomography CT (FDG PET-CT).

1.9.7 For people with smouldering myeloma and disease progression, consider one of the following (taking into consideration previous imaging tests):

- whole-body MRI
- whole-body low-dose CT
- whole-body CT
- spinal MRI
- FDG PET-CT.

1.10 Managing relapsed myeloma

First relapse

NICE has a suite of technology appraisal guidance on myeloma either published or in development. These published technology appraisals cover NICE's position in relation to primary disease treatment, salvage therapy for relapsed myeloma and consolidation/maintenance therapy after primary management. The recommendations in this guideline complement the existing technology appraisals, giving further guidance in addition to the technology appraisals where myeloma-related subgroups are not included.

1.10.1 Bortezomib monotherapy is recommended as an option for the treatment of progressive multiple myeloma in people who are at first relapse having received 1 prior therapy and who have undergone, or are unsuitable for, bone marrow transplantation, under the following circumstances:

- the response to bortezomib is measured using serum M protein after a maximum of 4 cycles of treatment, and treatment is continued only in people who have a complete or partial response (that is, reduction in serum M

protein of 50% or more or, where serum M protein is not measurable, an appropriate alternative biochemical measure of response), and

- the manufacturer rebates the full cost of bortezomib for people who, after a maximum of 4 cycles of treatment, have less than a partial response (as defined above). [This recommendation is from [NICE technology appraisal guidance on bortezomib monotherapy for relapsed multiple myeloma.](#)]

1.10.2 People currently receiving bortezomib monotherapy who do not meet the criteria in recommendation 1.10.1 should have the option to continue therapy until they and their clinicians consider it appropriate to stop. [This recommendation is from [NICE technology appraisal guidance on bortezomib monotherapy for relapsed multiple myeloma.](#)]

Second autologous stem cell transplantation

1.10.3 Offer a second autologous stem cell transplant, if it is suitable, to people with relapsed myeloma who have:

- completed re-induction therapy without disease progression and
- had a response duration of more than 24 months after their first autologous stem cell transplant.

1.10.4 Consider a second autologous stem cell transplant, if it is suitable, for people with relapsed myeloma who have:

- completed re-induction therapy without disease progression and
- had a response duration of between 12 and 24 months after their first autologous stem cell transplant.

1.10.5 Be aware that a second autologous stem cell transplant is more likely to be suitable for people with relapsed myeloma if they have:

- had a good response to the first autologous stem cell transplant
- a lower International Staging System (ISS) stage

- not had many prior treatments
- good overall fitness, based on resilience, frailty and performance status
- no adverse fluorescence in-situ hybridisation (FISH) results.

Subsequent therapy

- 1.10.6 Lenalidomide in combination with dexamethasone is recommended, within its licensed indication, as an option for the treatment of multiple myeloma only in people who have received two or more prior therapies, with the following condition:
- The drug cost of lenalidomide (excluding any related costs) for people who remain on treatment for more than 26 cycles (each of 28 days; normally a period of 2 years) will be met by the manufacturer. [This recommendation is from [NICE technology appraisal guidance on lenalidomide for the treatment of multiple myeloma in people who have received at least one prior therapy.](#)]
- 1.10.7 People currently receiving lenalidomide for the treatment of multiple myeloma, but who have not received 2 or more prior therapies, should have the option to continue therapy until they and their clinicians consider it appropriate to stop. [This recommendation is from [NICE technology appraisal guidance on lenalidomide for the treatment of multiple myeloma in people who have received at least one prior therapy.](#)]
- 1.10.8 Pomalidomide, in combination with dexamethasone, is not recommended within its marketing authorisation for treating relapsed and refractory multiple myeloma in adults who have had at least 2 previous treatments, including lenalidomide and bortezomib, and whose disease has progressed on the last therapy. [This recommendation is from [NICE technology appraisal guidance on pomalidomide for relapsed and refractory multiple myeloma previously treated with lenalidomide and bortezomib.](#)]
- 1.10.9 People whose treatment with pomalidomide was started within the NHS before this guidance was published should be able to continue treatment until they and their NHS clinician consider it appropriate to stop. [This recommendation is from

NICE technology appraisal guidance on pomalidomide for relapsed and refractory multiple myeloma previously treated with lenalidomide and bortezomib.]

Terms used in this guideline

Smouldering myeloma

In this guideline, only recommendations that specifically refer to smouldering myeloma apply to this condition.

Recommendations for research

The guideline committee has made the following recommendations for research. The committee's full set of research recommendations is detailed in the [full guideline](#).

1 Diagnostic investigations to predict treatment outcomes

What is the prognostic value of the Hevylite assay and ratio compared with other prognostic factors and tests, including the serum-free light-chain assay and fluorescence in-situ hybridisation (FISH), in people with newly diagnosed myeloma who are starting treatment?

Why this is important

Hevylite is a new assay that some studies have indicated is a useful prognostic tool. However, it is not clear how robustly it has been evaluated against other prognostic factors and tests, or whether it is an independent prognostic factor. The Hevylite assay should be evaluated in an accredited centralised laboratory independent of links with the manufacturer. Outcomes of interest are overall response, complete response, minimal residual disease, progression-free survival, overall survival and resource use.

2 Imaging investigations for newly diagnosed myeloma

What is the comparative effectiveness of whole-body MRI, fluorodeoxyglucose positron emission tomography CT (FDG PET-CT) and whole-body low-dose CT in detecting lesions that may determine the start of treatment for people with newly diagnosed myeloma?

Why this is important

Newer imaging techniques are replacing skeletal surveys for assessing myeloma-related bone disease in people with newly diagnosed myeloma. However, the most effective technique is not known. Outcomes of interest are lesion detection, sensitivity and

specificity for myeloma-related bone disease, patient acceptability, incremental upstaging, radiation exposure, risk of second primary cancer, the impact of additional information on predicting progression-free survival, overall survival and skeletal-related events.

3 Management of smouldering myeloma

Which combinations of FISH, molecular technologies, bone marrow plasma cell percentage, whole-body imaging, immunophenotype, serum-free light-chain levels or ratio, Hevylite, paraprotein levels, immunoparesis, and International Staging System (ISS) are most effective at risk stratification for people with smouldering myeloma?

What is the comparative effectiveness of fixed duration treatment (with or without bone-directed therapy), continuous treatment (with or without bone-directed therapy) and no treatment (with or without bone-directed therapy) for people with smouldering myeloma?

Why this is important

Changes to the International Myeloma Working Group definitions of smouldering myeloma and myeloma have affected the risk stratification process for smouldering myeloma. It is unclear if the previous risk stratification approach remains valid. It is also unclear if earlier treatment will be of benefit to people with smouldering myeloma. This study should be a randomised multi-centre prospective trial for patients with newly diagnosed smouldering myeloma (as defined by the International Myeloma Working Group 2014 classification). Outcomes of interest are time to biochemical and/or clinical progression, overall survival, adverse events, quality of life and resource use.

4 Allogeneic stem cell transplantation

What is the effectiveness of combined autologous–allogeneic stem cell transplantation compared with autologous stem cell transplantation, plus consolidation and maintenance treatment in chemosensitive patients at first response or first relapse?

Why this is important

There are conflicting data from a small number of studies on long-term survival following auto/allo stem cell transplantation compared with autologous stem cell transplantation.

These studies were performed before thalidomide, bortezomib and lenalidomide were used as myeloma treatments. These drugs produce better responses and also have the capacity to affect immunological responses after the transplant. Research is needed to see if there is a role for auto/allo stem cell transplant in the ongoing treatment of myeloma. Outcomes of interest are progression-free survival, overall survival, transplant-related mortality, quality of life, early and late toxicity including graft-versus-host-disease (GvHD) and resource use. This research should be included as an option in appropriate mainstream clinical trials for myeloma.

5 Bisphosphonates for the prevention of bone disease

What is the effectiveness of monthly zoledronic acid given indefinitely compared with zoledronic acid given for a fixed duration in patients with myeloma?

Why this is important

There is good-quality evidence to support the use of zoledronic acid to prevent bone disease in people with myeloma. However, the optimal frequency and duration of treatment is not clearly defined and needs further research, particularly given the quality-of-life implications for people needing regular, life-long visits to hospital. This study should be a randomised controlled trial. Outcomes of interest are skeletal-related events, progression-free survival, overall survival, utility of bone biomarkers, incidence of osteonecrosis of the jaw, quality of life and resource use.

Context

Because of the changes to the [International Myeloma Working Group definition of smouldering myeloma](#), it was not possible to make any recommendations for clinical practice on managing this condition. The new definition has changed how smouldering myeloma and myeloma are differentiated, and there is currently no evidence available that is using the new definitions.

This guideline covers adults (aged 16 years and over):

- who are referred to secondary care with suspected myeloma
- with newly diagnosed or relapsed myeloma (including high-risk myeloma and primary plasma cell leukaemia).

This guideline does not cover people who have:

- a solitary plasmacytoma without myeloma
- amyloid light-chain amyloidosis in the absence of myeloma
- paraproteins secondary to other conditions.

Finding more information and committee details

To find NICE guidance on related topics, including guidance in development, see the [NICE topic pages on blood and bone marrow cancers, complications of cancer, embolism and thrombosis](#) and [end of life care](#).

For full details of the evidence and the guideline committee's discussions, see the [full guideline](#). You can also find information about [how the guideline was developed](#), including details of the committee.

NICE has produced [tools and resources to help you put this guideline into practice](#). For general help and advice on putting our guidelines into practice, see [resources to help you put NICE guidance into practice](#).

Update information

October 2018: Recommendations on preventing thrombosis were replaced by [NICE's guideline on venous thromboembolism in over 16s](#).

Minor changes since publication

February 2026: We simplified the guideline by removing recommendations on general principles of care that are covered in other NICE guidelines.

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