NATIONAL INSTITUTE FOR HEALTH AND CARE EXCELLENCE

Health Technology Evaluation

**Elranatamab for treating relapsed or refractory multiple myeloma after 3 therapies**

**Final scope**

# Remit/evaluation objective

To appraise the clinical and cost effectiveness of elranatamab within its marketing authorisation for treating relapsed or refractory multiple myeloma after 3 therapies.

# Background

Multiple myeloma is a form of cancer that arises from plasma cells (a type of white blood cell) in the bone marrow. Myeloma cells produce large quantities of an abnormal antibody, known as paraprotein. Unlike normal antibodies, paraprotein has no useful function and lacks the capacity to fight infection. Myeloma cells supress the development of normal blood cells that are responsible for fighting infection (white blood cells), carrying oxygen around the body (red blood cells) and blood clotting (platelets). The term multiple myeloma refers to the presence of more than one site of affected bone at the time of diagnosis. People with multiple myeloma can experience bone pain, bone fractures, tiredness (due to anaemia), infections, hypercalcaemia (too much calcium in the blood) and kidney problems.

Approximately 5,000 people are diagnosed with multiple myeloma in England each year (2016 to 2018 data).1 Five-year prevalence of multiple myeloma in the UK is 26 per 100,000.2 It is most frequently diagnosed in older people, with 43% of new cases of multiple myeloma in England in people aged 75 years or older.1 The 5-year survival rate for adults with multiple myeloma in England and Wales is estimated to be 52%.3 Multiple myeloma is more common in men than in women.4 The incidence rates are also reported to be lower in the Asian ethnic group, higher in the Black ethnic group, and similar in people of mixed or multiple ethnicity, compared with the White ethnic group, in England (2013-2017 data).4

The main aims of therapy are to prolong survival and maintain a good quality of life by controlling the disease and relieving symptoms. If the disease progresses after initial treatment, the choice of subsequent therapy is influenced by previous treatment and response to it, duration of remission, comorbidities and patient preference.

For people who have had at least 2 prior therapies:

* [NICE technology appraisal guidance 171](https://www.nice.org.uk/guidance/TA171) recommends lenalidomide plus dexamethasone as a treatment option for people who have had at least 2 previous therapies.
* [NICE technology appraisal guidance 380](https://www.nice.org.uk/guidance/TA380) recommends panobinostat plus bortezomib and dexamethasone as a treatment option for adults who have had at least 2 previous therapies including bortezomib and an immunomodulatory agent.
* [NICE technology appraisal guidance 870](https://www.nice.org.uk/guidance/ta870) recommends ixazomib plus lenalidomide and dexamethasone as a treatment option for adults who have had 2 or 3 previous therapies.

For people who have had at least 3 prior therapies:

* [NICE technology appraisal guidance 427](https://www.nice.org.uk/guidance/TA427) recommends pomalidomide plus low-dose dexamethasone as a treatment option for adults who have had at least 3 previous treatments including both lenalidomide and bortezomib.
* [NICE technology appraisal guidance 783](https://www.nice.org.uk/guidance/ta783) recommends daratumumab monotherapy for use as a treatment option for adults who have had 3 previous therapies including a proteasome inhibitor and an immunomodulator.
* [NICE technology appraisal guidance 658](https://www.nice.org.uk/guidance/ta658) recommends isatuximab plus pomalidomide and dexamethasone for use within the Cancer Drugs Fund as a treatment option for adults who have had at least 3 previous treatments including both lenalidomide and a proteasome inhibitor.

# The technology

Elranatamab (brand name unknown, Pfizer) does not currently have a marketing authorisation in the UK for relapsed or refractory multiple myeloma after 3 prior therapies. It has been studied in a clinical trial in people with relapsed or refractory multiple myeloma who are refractory to at least one proteasome inhibitor, one immunomodulatory drug, and one anti-CD38 monoclonal antibody.

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| **Intervention(s)** | Elranatamab |
| **Population(s)** | People with relapsed or refractory multiple myeloma after at least 3 prior therapies |
| **Comparators** | * Lenalidomide plus dexamethasone * Panobinostat plus bortezomib and dexamethasone * Pomalidomide plus low-dose dexamethasone * Daratumumab monotherapy * Ixazomib plus lenalidomide and dexamethasone * Belantamab mafodotin (subject to NICE evaluation) * Cyclophosphamide plus dexamethasone |
| **Outcomes** | The outcome measures to be considered include:   * overall survival * progression-free survival * response rates * adverse effects of treatment * health-related quality of life. |
| **Economic analysis** | The reference case stipulates that the cost effectiveness of treatments should be expressed in terms of incremental cost per quality-adjusted life year.  If the technology is likely to provide similar or greater health benefits at similar or lower cost than technologies recommended in published NICE technology appraisal guidance for the same indication, a cost comparison may be carried out.  The reference case stipulates that the time horizon for estimating clinical and cost effectiveness should be sufficiently long to reflect any differences in costs or outcomes between the technologies being compared.  Costs will be considered from an NHS and Personal Social Services perspective.  The availability of any commercial arrangements for the intervention, comparator and subsequent treatment technologies will be taken into account.  The availability and cost of biosimilar and generic products should be taken into account. |
| **Other considerations** | Guidance will only be issued in accordance with the marketing authorisation. Where the wording of the therapeutic indication does not include specific treatment combinations, guidance will be issued only in the context of the evidence that has underpinned the marketing authorisation granted by the regulator. |
| **Related NICE recommendations** | **Related Technology Appraisals:**  [‘Daratumumab monotherapy for treating relapsed and refractory multiple myeloma’](https://www.nice.org.uk/guidance/ta783) (2022) NICE technology appraisal guidance 783. Review date 2025.  ‘[Isatuximab with pomalidomide and dexamethasone for treating relapsed and refractory multiple myeloma](https://www.nice.org.uk/Guidance/TA658)’ (2020) NICE technology appraisal guidance 658. Review date to be confirmed.  [‘Lenalidomide for the treatment of multiple myeloma in people who have received at least 2 prior therapies’](https://www.nice.org.uk/guidance/ta171) (2019) NICE technology appraisal guidance 171. No current plans to review this guidance.  ‘[Pomalidomide for multiple myeloma previously treated with lenalidomide and bortezomib](https://www.nice.org.uk/Guidance/TA427)’ (2017) NICE technology appraisal guidance 427. Review date to be confirmed.  [‘Panobinostat for treating multiple myeloma after at least 2 previous treatments’](https://www.nice.org.uk/guidance/TA380) (2016) NICE technology appraisal guidance 380. No current plans to review this guidance.  ‘[Ixazomib with lenalidomide and dexamethasone for treating relapsed or refractory multiple myeloma (CDF review of TA505)](https://www.nice.org.uk/guidance/indevelopment/gid-ta10781)’ (2023) NICE technology appraisal guidance 870. No current plans to review this guidance.  **Related appraisals in development:**  [Belantamab mafodotin for treating relapsed or refractory multiple myeloma after 2 therapies](https://www.nice.org.uk/guidance/awaiting-development/gid-ta11018) NICE technology appraisal guidance [ID5108]. Publication date to be confirmed.  [‘Belantamab mafodotin for treating relapsed or refractory multiple myeloma after 4 or more therapies’](https://www.nice.org.uk/guidance/indevelopment/gid-ta10568) NICE technology appraisal guidance [ID2701]. Publication expected August 2023.  ‘[Ciltacabtagene autoleucel for treating relapsed and lenalidomide-refractory multiple myeloma after 1 to 3 therapies](https://www.nice.org.uk/guidance/awaiting-development/gid-ta10905)’ NICE technology appraisal guidance [ID4012]. Publication date to be confirmed.  ‘[Idecabtagene vicleucel for treating relapsed and refractory multiple myeloma in people who have received at least 3 prior therapies](https://www.nice.org.uk/guidance/indevelopment/gid-ta10672)’ NICE technology appraisal guidance [ID1442]. Publication date to be confirmed.  ‘[Isatuximab with pomalidomide and dexamethasone for treating relapsed and refractory multiple myeloma [Review of TA658]](https://www.nice.org.uk/guidance/indevelopment/gid-ta10979)’ NICE technology appraisal guidance [ID4067]. Publication date to be confirmed.  ‘[Selinexor with bortezomib and low-dose dexamethasone for treating relapsed refractory multiple myeloma](https://www.nice.org.uk/guidance/indevelopment/gid-ta10646)’ NICE technology appraisal guidance [ID3797]. Publication date to be confirmed.  **Related Guidelines:**  [‘Myeloma: diagnosis and management’](https://www.nice.org.uk/guidance/ng35) (2018). NICE guideline 35. No current plans to review this guidance.  ‘[Haematological cancers: improving outcomes](https://www.nice.org.uk/guidance/ng47)’ (2016). NICE guidance 47. No current plans to review this guidance.  **Related Quality Standards:**  ‘[Haematological cancers](https://www.nice.org.uk/guidance/qs150)’ (2017) NICE quality standard 150. No current plans to review this guidance. |
| **Related National Policy** | The NHS Long Term Plan, 2019. [NHS Long Term Plan](https://www.longtermplan.nhs.uk/publication/nhs-long-term-plan/)  NHS England (2018/2019) [NHS manual for prescribed specialist services (2018/2019)](https://www.england.nhs.uk/publication/manual-for-prescribed-specialised-services/)  NHS England (2020) [Bendamustine for relapsed multiple myeloma (all ages).](https://www.england.nhs.uk/wp-content/uploads/2020/07/1608-Policy-bendamustine-relapsed-multiple-myeloma.pdf) Clinical Commissioning Policy. Reference: 200604P |

# References

1. Cancer Research UK, [Myeloma incidence statistics](https://www.cancerresearchuk.org/health-professional/cancer-statistics/statistics-by-cancer-type/myeloma/incidence#heading-Zero). Accessed October 2022.
2. United Kingdom Fact sheet, [International Agency for Research on Cancer](https://gco.iarc.fr/today/data/factsheets/populations/826-united-kingdom-fact-sheets.pdf). Accessed October 2022.
3. Cancer Research UK, [Myeloma survival statistics](https://www.cancerresearchuk.org/health-professional/cancer-statistics/statistics-by-cancer-type/myeloma/survival#heading-Zero). Accessed October 2022.
4. Cancer Research UK, [Myeloma statistics](https://www.cancerresearchuk.org/health-professional/cancer-statistics/statistics-by-cancer-type/myeloma#heading-Zero). Accessed October 2022.