

Osimertinib for untreated EGFR mutation-positive non-small-cell lung cancer

Technology appraisal guidance

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Your responsibility

The recommendations in this guidance represent the view of NICE, arrived at after careful consideration of the evidence available. When exercising their judgement, health professionals are expected to take this guidance fully into account, alongside the individual needs, preferences and values of their patients. The application of the recommendations in this guidance is at the discretion of health professionals and their individual patients and do not override the responsibility of healthcare professionals to make decisions appropriate to the circumstances of the individual patient, in consultation with the patient and/or their carer 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](#).

Commissioners and/or providers have a responsibility to provide the funding required to enable the guidance to be applied when individual health professionals and their patients wish to use it, in accordance with the NHS Constitution. They should do so 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.

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.

Contents

1 Recommendation	4
2 Information about osimertinib	5
Marketing authorisation indication	5
Dosage in the marketing authorisation	5
Price	5
3 Committee discussion	6
Clinical need	6
Clinical evidence	7
Modelling of overall survival	9
Cost-effectiveness estimate	12
End of life	13
Innovation	17
Other factors	17
Conclusion	17
4 Implementation	18
5 Appraisal committee members and NICE project team	20
Appraisal committee members	20
NICE project team	20

This guidance replaces TA621.

1 Recommendation

- 1.1 Osimertinib is recommended, within its marketing authorisation, as an option for untreated locally advanced or metastatic epidermal growth factor receptor (EGFR) mutation-positive non-small-cell lung cancer (NSCLC) in adults. It is recommended only if the company provides osimertinib according to the [commercial arrangement](#).

Why the committee made this recommendation

Locally advanced or metastatic EGFR mutation-positive NSCLC is usually first treated with afatinib, erlotinib or gefitinib.

Evidence from a randomised controlled trial suggests that people who take osimertinib live longer than people who take erlotinib or gefitinib. They also live longer before their disease gets worse. There is some uncertainty about the comparison of osimertinib with afatinib, which may be more effective than erlotinib and gefitinib, because there is no direct evidence comparing them.

But, because of a new commercial arrangement, the cost-effectiveness estimates for osimertinib are within what NICE considers an acceptable use of NHS resources. So, osimertinib is recommended.

2 Information about osimertinib

Marketing authorisation indication

- 2.1 Osimertinib (Tagrisso, AstraZeneca) is indicated 'for the first-line treatment of adult patients with locally advanced or metastatic non-small cell lung cancer (NSCLC) with activating epidermal growth factor receptor (EGFR) mutations'.

Dosage in the marketing authorisation

- 2.2 The dosage schedule is available in the [summary of product characteristics](#).

Price

- 2.3 The price is £5,770 for 80 mg and 40 mg osimertinib (pack of 30 tablets, excluding VAT; BNF online, accessed August 2020). The company has a commercial arrangement that makes osimertinib available to the NHS with a discount. The size of the discount is commercial in confidence. It is the company's responsibility to let relevant NHS organisations know details of the discount.

3 Committee discussion

The [appraisal committee](#) considered evidence submitted by AstraZeneca as part of the previous guidance (NICE technology appraisal guidance 621), another submission by AstraZeneca for the rapid review, reviews of these submissions by the evidence review group (ERG), and the technical report developed as part of the previous guidance through engagement with stakeholders. See the [committee papers](#) for full details of the evidence.

The appraisal committee was aware that several issues had been resolved during the technical engagement stage, and agreed that:

- The utility value of 0.678 (from the AURA 2 trial, second-line treatment with osimertinib) was more representative of people in the progressed disease state (table 3, page 24 to 25 of the technical report).
- A combined approach to determine the appropriate resource costs for people in the progressed disease state was acceptable (table 3, pages 24 to 25 of the technical report).

It recognised that there were remaining areas of uncertainty associated with the analyses presented (table 2, pages 22 to 23 of the technical report), and took these into account in its decision making. As part of the previous guidance, it discussed the following issues, which were outstanding after the technical engagement stage.

Clinical need

People would welcome a new treatment option

- 3.1 The patient experts explained that people with untreated locally advanced or metastatic epidermal growth factor receptor (EGFR) mutation-positive non-small-cell lung cancer (NSCLC) are often very unwell, with many distressing symptoms. There are no curative treatments. The prognosis is generally poor despite treatments such as targeted therapies and immunotherapy. People would therefore welcome new treatments that improve their symptoms and quality of life, and increase how long they live (even if this increase is only small). Locally

advanced or metastatic EGFR mutation-positive NSCLC is first treated with an EGFR tyrosine kinase inhibitor, such as afatinib, gefitinib or erlotinib, in line with [NICE guidance on afatinib, gefitinib and erlotinib](#). The clinical experts explained that people would usually be offered afatinib, based on the clinical evidence (see [section 3.4](#)). After afatinib, gefitinib or erlotinib, people may be offered either osimertinib if they have developed the T790M resistance mutation in the EGFR gene (in line with [NICE's technology appraisal guidance on osimertinib for treating EGFR T790M mutation-positive advanced NSCLC](#)), or chemotherapy if not. People who are not well enough to have further treatment would be offered best supportive care. After chemotherapy, people may be offered immunotherapy, docetaxel with or without nintedanib, or best supportive care. The clinical experts stated that osimertinib would be beneficial as an additional treatment option because it is better tolerated than existing treatments, with fewer side effects. Also, if osimertinib was a first-line treatment option, it would remove the need for T790M mutation testing before second-line treatment. This involves a biopsy, which is invasive and can be psychologically distressing. The committee agreed that additional options would be beneficial and concluded that osimertinib would be a useful addition to first-line treatment.

Clinical evidence

The FLAURA trial is broadly generalisable to people with untreated locally advanced or metastatic EGFR mutation-positive NSCLC in England

- 3.2 The clinical evidence for osimertinib came from the FLAURA randomised controlled trial. FLAURA compared the efficacy and safety of osimertinib with standard care (erlotinib or gefitinib) for people with locally advanced or metastatic EGFR mutation-positive NSCLC. Patients in the trial had either the exon 19 deletion (del19) or exon 21 (L858R) EGFR mutation. The clinical experts explained that these 2 mutations account for around 90% of all EGFR mutations. Also, most trials only include people with these mutations, including the trials that were carried out with other tyrosine kinase inhibitors. The committee acknowledged that, although other mutations may not respond as well to osimertinib, the marketing

authorisation indication is not restricted to these 2 mutations (see [section 2](#)). It therefore agreed that the EGFR mutation status of patients in FLAURA generally reflected that seen in NHS clinical practice in England. The inclusion criteria allowed people with stable brain metastases to enter the trial but limited the trial population to people with an Eastern Cooperative Oncology Group (ECOG) performance score of 0 or 1. For this reason, the committee was aware that the clinical trial population may be in better health than people with stage 3b or stage 4 NSCLC in the NHS and that people with many comorbidities may not have been included in the trial. Also, it noted that afatinib was not a comparator in the standard care arm in FLAURA (see [section 3.4](#)) and that many subsequent treatments used in the trial are not routinely used in the NHS. Despite these concerns, the clinical experts explained that the evidence from FLAURA was broadly generalisable to NHS clinical practice. The committee agreed with the clinical experts.

Osimertinib extends progression-free and overall survival compared with gefitinib and erlotinib but the size of the benefit is unclear

- 3.3 An interim analysis of FLAURA showed that progression-free survival was statistically significantly longer with osimertinib than with erlotinib or gefitinib. At the interim data cut (12 June 2017; presented in the previous guidance), median progression-free survival was 18.9 months for osimertinib (95% confidence interval [CI] 15.2 to 21.4) and 10.2 months for standard care (95% CI 9.6 to 11.1). The hazard ratio was 0.46 (95% CI 0.37 to 0.57; $p < 0.001$). Overall survival data were very immature (25% of events) but the interim results showed that osimertinib extended overall survival compared with standard care. This produced a hazard ratio of 0.63 (95% CI 0.45 to 0.88; $p = 0.007$), which was not statistically significant (a p value of less than 0.0015 was needed for the result to be significant). The committee acknowledged that the FLAURA data were very immature and that there was uncertainty in overall survival because of the number of events still to be reported. The trial reported more data on overall survival after the previous guidance was published, but submitting this evidence was beyond the scope of the rapid review process, so it could not be fully considered by the committee. Therefore the committee concluded, based on the evidence for the previous

guidance, that osimertinib lengthened progression-free survival, and possibly overall survival, compared with erlotinib or gefitinib. But it also concluded that the overall survival benefit was difficult to establish because the data were very immature.

EGFR tyrosine kinase inhibitors do not all have equal efficacy

3.4 The relevant comparators for this technology appraisal are erlotinib, gefitinib and afatinib. FLAURA compared osimertinib with either gefitinib or erlotinib, but not with afatinib. The Cancer Drugs Fund clinical lead noted that afatinib is currently the most prescribed EGFR tyrosine kinase inhibitor in England for this population. He also stated that previous trials, such as LUX-Lung 7, showed that afatinib statistically significantly improved progression-free survival compared with gefitinib. The clinical experts agreed that gefitinib and erlotinib are likely to have equal efficacy. They stated that people taking afatinib had a better response rate to treatment, a longer duration of response and longer progression-free survival than with erlotinib and gefitinib. Also, they usually stayed on afatinib for longer. The company stated that LUX-Lung 7 did not show a statistically significant increase in overall survival for afatinib compared with gefitinib. It therefore assumed that afatinib was equivalent in efficacy to erlotinib and gefitinib in its economic model. However, the clinical experts explained that LUX-Lung 7 was not powered (that is, it did not have enough people in the trial) to show a difference in overall survival compared with gefitinib. The ERG did its own exploratory indirect treatment comparison that suggested osimertinib statistically significantly improved progression-free survival compared with afatinib but showed no statistically significant difference in overall survival. The committee concluded that there was evidence of improved progression-free survival with afatinib compared with gefitinib, and erlotinib and gefitinib cannot be assumed to have equal efficacy with afatinib.

Modelling of overall survival

Assuming a 6-year treatment benefit for osimertinib is optimistic

3.5 The company used a partitioned survival structure with 3 health states

(progression-free, progressed disease and death) to model overall survival in FLAURA. It used a time horizon of 20 years to capture all relevant costs and benefits for people having treatment. The company initially assumed a treatment benefit for osimertinib for the full 20-year period. The committee agreed with the ERG and clinical experts that this assumption was optimistic considering the data available and would have to be adjusted to reflect a more realistic benefit from osimertinib treatment. The company therefore revised its base case, assuming a 6-year duration of treatment effect after the start of treatment (that is, applying a hazard ratio of 1 to both the osimertinib and standard care arms 6 years after starting treatment). The committee recalled that, in previous appraisals for locally advanced or metastatic NSCLC, the preferred treatment effect duration for immunotherapies was 3 years to 5 years. However, it acknowledged that these appraisals involved drugs with a different mechanism of action to osimertinib and a maximum treatment duration. Therefore, it was not appropriate to compare them. The clinical experts agreed that, because osimertinib is associated with improved progression-free survival and duration of response, treatment effect would continue after symptomatic and radiological progression for some people. They stated that this could plausibly give about 3 months of additional benefit after stopping treatment with osimertinib compared with erlotinib and gefitinib. The clinical experts believed that because osimertinib penetrates the blood–brain barrier better than erlotinib and gefitinib, it may help improve control of brain metastases. The committee recalled that afatinib yields longer progression-free survival than erlotinib and gefitinib, but because there was no direct evidence comparing osimertinib with afatinib (see [section 3.4](#)), it could not establish how osimertinib compared with afatinib in terms of progression-free and overall survival. The ERG's preferred analyses used durations of 3 years and 5 years. The ERG explained that the company's 6-year duration of treatment effect would mean that people who stopped taking osimertinib within 1 year or 2 years of starting it would still benefit for the full 6 years. The ERG emphasised the limitations of modelling the duration of treatment effect with a partitioned survival model. This is because a crude approach is needed to make adjustments around the assumptions (for example, assuming equivalence at a single time point). The ERG noted that this does not fully reflect what happens in a clinical setting. The committee concluded that a 6-year duration of treatment effect for

osimertinib was optimistic and that, without more evidence, the ERG's analysis using a 3-year or 5-year duration of treatment effect was more appropriate.

The economic model does not capture the benefits of subsequent treatments appropriately

- 3.6 The committee was aware of [NICE's position statement on handling comparators and treatment sequences in the Cancer Drugs Fund](#). This states that 'products recommended for use in the Cancer Drugs Fund after 1 April 2016 should not be considered as comparators, or appropriately included in a treatment sequence, in subsequent relevant appraisals'. But the committee accepted that it could consider the company's approach of including osimertinib as a subsequent treatment (in line with [NICE's guidance on osimertinib for treating locally advanced or metastatic EGFR T790M mutation-positive NSCLC](#)) in the model in this appraisal, because this reflected that some patients in FLAURA did receive osimertinib as a subsequent treatment. People in the standard care arm in the model could have osimertinib as a second-line treatment, assumed to be 33% of people (based on clinical opinion). The committee noted that, although the costs of osimertinib as a second-line treatment were applied in the standard care arm of the model, efficacy was not fully captured given that only around 20% of people had osimertinib as a second-line treatment in the trial at the time of the interim analysis. The committee was aware that the subsequent treatments used in the trial may not reflect NHS practice. It noted that different subsequent therapies would mean different survival prospects and health states that cannot be captured in the modelling. The ERG explained that, to overcome the limitations of the model in capturing the efficacy of subsequent treatments, and to create more flexibility to explore varying the duration of treatment benefits, additional health states would be needed. The ERG described how an individual patient simulation model would better account for these issues but it would need lots of additional data, and trial data are usually immature. The committee agreed that the company's model was broadly appropriate for decision making. It acknowledged the limitations of the model and, given the immaturity of the data, concluded that the model did not fully capture the benefits of subsequent treatments appropriately.

The company's modelling of overall survival is appropriate, but the immaturity of the data introduces uncertainty in estimating overall survival

3.7 At the interim data cut considered in the previous guidance, median overall survival was not achieved in either the osimertinib or standard care arm. To estimate the overall survival of people in FLAURA, the company used a piecewise Weibull extrapolation of the Kaplan–Meier curve, which was based on observed data up to 7.90 months in the trial. This estimated that mean overall survival was 66.96 months with osimertinib and 44.39 months with standard care, assuming a 20-year time horizon in the model. The committee was aware that this extrapolation resulted in the most conservative piecewise survival estimates of those presented and fitted the data well. It understood that the FLAURA data were immature (only 25% of events occurring), which introduced uncertainty into the survival estimates, and that further data collection is planned. It concluded that, although the company's and ERG's preferred choice of distribution for modelling overall survival was appropriate, the immaturity of the data introduces uncertainty in estimating the results.

Cost-effectiveness estimate

The most plausible ICER for osimertinib is within the range NICE normally considers a cost-effective use of NHS resources

- 3.8 The committee considered the additional analyses presented for the rapid review, which incorporated the updated confidential commercial arrangement for osimertinib, and its preferred modelling assumptions:
- A treatment effect duration (that is, from the start of treatment) of 3 years to 5 years (see [section 3.5](#)).
 - Weibull extrapolation of overall survival in both the osimertinib and standard care arms (see [section 3.7](#)).
 - A utility value of 0.678 (see table 3, pages 24 to 25 of the technical report).

These assumptions were used in a fully incremental analysis (calculating incremental quality-adjusted life year [QALY] gains and costs along a list of treatment options ranked by ascending cost), which incorporated all of the commercial arrangements for osimertinib and the comparators (gefitinib, erlotinib and afatinib). The committee was aware that NHS England considered that the commercial arrangement delivered additional value and included additional commercial information from NHS England in its decision making. Taking into account all the commercial considerations, the incremental cost-effectiveness ratio (ICER) was within the range NICE normally considers a cost-effective use of NHS resources. The exact ICERs cannot be reported here because they are commercial in confidence. The committee acknowledged that, given the available evidence from LUX-Lung 7 and clinical expert opinion, it is possible that afatinib has better efficacy than gefitinib and erlotinib (see [section 3.4](#)) and factored this into its decision making. It agreed that its most plausible ICER based on the evidence presented was within the range that NICE considers an effective use of NHS resources.

End of life

Osimertinib is likely to extend life by over 3 months

- 3.9 The committee considered the advice about life-extending treatments for people with a short life expectancy in [NICE's guide to the methods of technology appraisal](#). It recalled that, at the interim data cut presented and considered in the previous guidance, median overall survival had not been reached in FLAURA, and that the increased survival in the osimertinib arm was not statistically significantly different from the standard care arm (see [section 3.3](#)). However, it also noted that the company's economic model predicted mean overall survival would be 22 months longer with osimertinib than with standard care. Based on evidence from FLAURA and predictions from the economic model (using the committee's preferred assumptions), the committee concluded that osimertinib was likely to extend life by over 3 months and therefore met the extension-to-life criterion.

FLAURA should be the primary data source for deciding if osimertinib meets the short life expectancy criterion

- 3.10 For the previous guidance, the company presented registry evidence from a real-world data source (National Cancer Registration and Analysis Service [NCRAS] Public Health England data between 2014 and 2016), showing that median overall survival for the population in England was less than 24 months. The committee recognised that there was potential value in real-world evidence from the NHS in England to help inform its decision making. However, it considered there were several reasons why it was not appropriate to use these as the primary data source in isolation for its decision making on the short life expectancy criterion:
- The committee recalled its conclusion that FLAURA was generalisable to clinical practice in England (see [section 3.2](#)). In addition, it noted the Cancer Drugs Fund clinical lead and ERG statements that it was inconsistent to use the FLAURA data to determine the overall survival benefit of osimertinib, but real-world evidence to determine life expectancy for people having standard care, without making some adjustments in the economic model (such as amending the efficacy estimates).
 - The committee noted that afatinib (the currently preferred and most prescribed EGFR tyrosine kinase inhibitor in England for this population) was not available for most of the time the NCRAS data were collected.
 - The committee recalled that after consultation on the technical report, the clinical experts stated that about 60% of people in clinical practice were alive 2 years after starting treatment with an EGFR tyrosine kinase inhibitor.
 - In the standard care arm, which used FLAURA data to inform the extrapolation of overall survival, the economic model (using the committee's preferred assumptions) predicted a median overall survival of 31.54 months and a mean overall survival of 44.39 months.
 - The committee was also aware that evidence from studies in similar populations, such as LUX-Lung 7 and ARCHER 1050, showed that median overall survival was more than 24 months.
 - The committee noted that the registry data were difficult to compare directly with the FLAURA data because possible confounders in the real-world

population (such as comorbidities) were not taken into account.

For these reasons, the committee considered it more appropriate to base its decision about estimating life expectancy on the FLAURA data, which the company had used in the economic model. The committee concluded that osimertinib did not meet the short life expectancy criterion of the end of life criteria.

Subgroup analyses from FLAURA do not show that osimertinib meets the criteria for life-extending treatment at the end of life

3.11 At consultation on the previous guidance, the company provided new analyses to support its case for meeting the end of life criteria when using the committee-preferred dataset (FLAURA). The company stated that the subgroup analyses most closely reflected the cohort of NHS patients in England. However, the Cancer Drugs Fund clinical lead's nominated deputy advised caution in interpreting these registry data because the systemic anticancer therapy (SACT) data set is currently incomplete and that gaps in the evidence exist between the secondary uses service data and the corresponding SACT data. The 3 subgroups were:

- No subsequent treatment with an EGFR tyrosine kinase inhibitor (because the company stated that people in England would not have a second EGFR tyrosine kinase inhibitor).
- An ECOG performance score of 1 (because people in England with this condition are usually less well than those in FLAURA).
- Non-Asian family origin (because most people with this condition in England are of non-Asian family origin and the company stated that they have poorer survival outcomes than people of Asian family origin).

The committee noted that the subgroups in the new analyses had not been combined to calculate a value for mean or median overall survival. It agreed that this added substantial uncertainty to interpreting the results because each subgroup was linked to a single characteristic and it was not possible to determine the degree of overlap between these groups. The committee noted that the company's overall survival estimates for all 3 subgroups in the

standard care arm were longer than 24 months (the modelled outputs are academic in confidence and cannot be reported). The ERG stated that because it did not have access to the FLAURA data, it could not confirm any of the company's results for the new analyses. The committee heard concerns regarding the relevance of the analyses:

- The Cancer Drugs Fund clinical lead's nominated deputy commented that many people do have more than 2 subsequent therapies after progression on an EGFR tyrosine kinase inhibitor.

The ERG noted that the overall survival for patients of non-Asian family origin may be shorter, but the results of the subgroup analysis were very similar to those of the intention-to-treat population in FLAURA and that no statistical testing of the difference was done. The committee agreed there was no conclusive evidence that ethnicity has an influence on overall survival and that other factors may be more influential. It noted that for [NICE's technology appraisal guidance on afatinib](#), the clinical experts stated that differences in the effectiveness of afatinib in NSCLC are more likely to be determined by EGFR mutation status than ethnicity.

The committee concluded the subgroup analyses presented in response to consultation on the previous guidance did not show that osimertinib meets the short life expectancy criterion. It noted that the short life expectancy criterion in the methods guide states 'normally less than 24 months' and discussed whether any flexibilities should be applied. The committee concluded that there were no exceptional circumstances that demanded additional flexibility in applying the end of life criteria (such as to ensure continued access to a highly effective treatment option that was perceived by patients and clinicians to be standard of care, in circumstances where access had been enabled years ahead of NICE publishing any guidance on the technology). The committee also concluded that, although the company's economic model suggests that the overall survival gain may potentially be high (see [section 3.7](#)), the immaturity of the trial data means there is considerable uncertainty about the magnitude of the benefit.

Innovation

Osimertinib may be innovative

- 3.12 The Cancer Drugs Fund clinical lead highlighted that follow up in FLAURA is short, so the economic model was unlikely to fully capture osimertinib's beneficial effect in the brain. They also stated that osimertinib is better tolerated than other EGFR tyrosine kinase inhibitors with respect to chronic grade 1 and grade 2 skin-related toxicities, and this benefit was not captured in the economic model. The committee also understood that having osimertinib for untreated EGFR mutation-positive NSCLC will reduce the need for repeat bronchoscopic biopsies in people to identify those eligible for osimertinib after an EGFR tyrosine kinase inhibitor (this is currently available via the Cancer Drugs Fund). The committee concluded that osimertinib may be innovative and agreed that there may be benefits that are not captured by the cost-effectiveness analyses.

Other factors

- 3.13 No equality or social value judgement issues were identified.

Conclusion

Osimertinib is recommended for routine use in the NHS

- 3.14 The committee concluded that the estimates of cost effectiveness for osimertinib were within the range that is considered to be a cost-effective use of NHS resources. This conclusion took into account the clinical evidence from FLAURA, all the relevant commercial arrangements, including an updated commercial arrangement for osimertinib and commercial information from NHS England, and the potential benefits of osimertinib not captured by the economic model. So, osimertinib is recommended for use in the NHS for untreated locally advanced or metastatic EGFR mutation-positive NSCLC in adults.

4 Implementation

- 4.1 Section 7(6) of the National Institute for Health and Care Excellence (Constitution and Functions) and the Health and Social Care Information Centre (Functions) Regulations 2013 requires clinical commissioning groups, NHS England and, with respect to their public health functions, local authorities to comply with the recommendations in this appraisal within 3 months of its date of publication.
- 4.2 Chapter 2 of Appraisal and funding of cancer drugs from July 2016 (including the new Cancer Drugs Fund) – a new deal for patients, taxpayers and industry states that for those drugs with a draft recommendation for routine commissioning, interim funding will be available (from the overall Cancer Drugs Fund budget) from the point of marketing authorisation, or from release of positive draft guidance, whichever is later. Interim funding will end 90 days after positive final guidance is published (or 30 days in the case of drugs with an Early Access to Medicines Scheme designation or fast track appraisal), at which point funding will switch to routine commissioning budgets. The NHS England and NHS Improvement Cancer Drugs Fund list provides up-to-date information on all cancer treatments recommended by NICE since 2016. This includes whether they have received a marketing authorisation and been launched in the UK.
- 4.3 The Welsh ministers have issued directions to the NHS in Wales on implementing NICE technology appraisal guidance. When a NICE technology appraisal recommends the use of a drug or treatment, or other technology, the NHS in Wales must usually provide funding and resources for it within 2 months of the first publication of the final appraisal document.
- 4.4 When NICE recommends a treatment 'as an option', the NHS must make sure it is available within the period set out in the paragraphs above. This means that, if a patient has untreated locally advanced or metastatic epidermal growth factor receptor (EGFR) mutation-positive non-small-cell lung cancer (NSCLC), and the doctor responsible for their care thinks that osimertinib is the right treatment, it should be available for use, in

line with NICE's recommendations.

5 Appraisal committee members and NICE project team

Appraisal committee members

The 4 technology appraisal committees are standing advisory committees of NICE. This topic was considered by [committee D](#).

Committee members are asked to declare any interests in the technology to be appraised. If it is considered there is a conflict of interest, the member is excluded from participating further in that appraisal.

The [minutes of each appraisal committee meeting](#), which include the names of the members who attended and their declarations of interests, are posted on the NICE website.

NICE project team

Each technology appraisal is assigned to a team consisting of 1 or more health technology analysts (who act as technical leads for the appraisal), a technical adviser and a project manager.

Thomas Paling and Stephen Robinson

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