NATIONAL INSTITUTE FOR HEALTH AND CARE EXCELLENCE

Final draft guidance

Dabrafenib with trametinib for treating BRAF V600E mutation-positive glioma in children and young people aged 1 year and over

1 Recommendations

- 1.1 Dabrafenib with trametinib is recommended, within its marketing authorisation, as an option for treating:
 - low-grade glioma (LGG) with a BRAF V600E mutation in children and young people aged 1 year and over who need systemic treatment
 - high-grade glioma (HGG) with a BRAF V600E mutation in children and young people aged 1 year and over after at least 1 radiation or chemotherapy treatment.

Dabrafenib with trametinib is only recommended if the company provides it according to the commercial arrangement (see <u>section 2</u>).

Why the committee made these recommendations

Glioma is a type of brain cancer that is classified into LGG or HGG based on how fast it grows. Usual treatment for glioma includes surgery, radiotherapy, chemotherapy and best supportive care.

In LGG, dabrafenib plus trametinib has been directly compared with chemotherapy in a clinical trial in people aged 1 to 17 years. It shows that people who have dabrafenib plus trametinib have longer before their condition gets worse than people who have chemotherapy.

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In HGG, dabrafenib plus trametinib has not been directly compared with any treatment. But, indirect comparisons suggest that people aged 1 to 17 years who have dabrafenib plus trametinib have longer before their condition gets worse than people who have chemotherapy or best supportive care.

When considering the condition's severity, and its effect on quality and length of life, the most likely cost-effectiveness estimates are within the range that NICE considers an acceptable use of NHS resources. So, dabrafenib plus trametinib is recommended.

2 Information about dabrafenib with trametinib

Marketing authorisation indication

- 2.1 Dabrafenib (Finlee, Novartis) in combination with trametinib (Spexotras, Novartis) is indicated for:
 - 'the treatment of paediatric patients aged 1 year and older with lowgrade glioma (LGG) with a BRAF V600E mutation who require systemic therapy'
 - 'the treatment of paediatric patients aged 1 year and older with highgrade glioma (HGG) with a BRAF V600E mutation who have received at least one prior radiation and/or chemotherapy treatment'.

Dosage in the marketing authorisation

2.2 The dosage schedule is available in the <u>summary of product</u> <u>characteristics for dabrafenib</u> and <u>trametinib</u>.

Price

2.3 The list price for dabrafenib is £2,800 per 420-pack of 10 mg dispersible tablets (company submission). The list price for trametinib is £376 per 4.7 mg bottle of 0.05 mg per ml powder for oral solution (company submission).

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2.4 The company has a commercial arrangement for each medicine (simple discount patient access scheme). This makes dabrafenib plus trametinib 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 <u>evaluation committee</u> considered evidence submitted by Novartis, a review of this submission by the external assessment group (EAG), and responses from stakeholders. See the <u>committee papers</u> for full details of the evidence.

The condition

BRAF V600E mutation-positive glioma

3.1 Gliomas are the most common type of brain cancer in children and young people. They develop from the glial cells that support the nerve cells of the brain and spinal cord. Gliomas are classified by how quickly they grow. Most gliomas are grade 1 or 2, referred to as low-grade glioma (LGG), and do not grow or only grow slowly. Grade 3 and 4 gliomas, referred to as high-grade glioma (HGG), grow rapidly. Consequently, HGG is associated with worse outcomes than LGG. BRAF is a gene that encodes the protein B-Raf, which influences cell growth. People with BRAF V600E mutation-positive LGG live for less time than people with glioma without the mutation. The patient experts emphasised the traumatic nature of a glioma diagnosis for children and young people, their families and caregivers, and the limitations of current treatment. They noted that glioma and its treatment can delay education, restrict socialising, and cause lasting emotional impact. They highlighted that the toxicity associated with conventional chemotherapy can lead to people with glioma and their caregivers choosing to stop treatment. The patient experts also explained that currently available treatments need regular travel to hospital, incur significant costs and need a substantial time

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commitment. For these reasons, people with glioma and their caregivers would value additional treatment options, particularly those that can be taken at home. The committee understood the comments from the patient experts about the effect of glioma on people who have it, their families and caregivers, and recognised that there is a high burden for people with LGG and HGG.

Clinical management

Treatment options

3.2 The clinical experts explained that the aims of treatment for glioma include stopping or delaying progression and improving neurological function and guality of life. Current treatment for LGG includes maximal surgical resection, followed by systemic chemotherapy or radiotherapy at first relapse or progression. At further relapse or progression, other chemotherapy regimens may be used. Current treatment for HGG includes maximal surgical resection followed by chemoradiotherapy (typically with adjuvant temozolomide – see NICE's technology appraisal guidance on carmustine implants and temozolomide for the treatment of newly diagnosed high-grade glioma). At relapse or progression, some people will have further chemotherapy, radiotherapy or resection and some people will have best supportive care. The clinical experts noted that there is an unmet need for treatments for children and young people with BRAF V600E mutation-positive glioma. The clinical experts thought that dabrafenib plus trametinib, as a treatment that is specific for BRAF V600E mutations, represents a step-change in care compared with current treatment. The committee concluded that current treatment for glioma is limited and that dabrafenib plus trametinib offers a new treatment option.

Comparators

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LGG cohort

3.3 The final NICE scope listed chemotherapy (including, but not limited to vincristine plus carboplatin) as the comparator for LGG. The company considered that vincristine plus carboplatin is the most relevant comparator available in LGG. It cited the Children's Cancer and Leukaemia Group (CCLG) guidelines that state vincristine plus carboplatin should be considered as first-line treatment for people with nonneurofibromatosis type 1 LGG. At second line, the CCLG guidelines recommend vinblastine monotherapy. For people with neurofibromatosis type 1 LGG, the guidelines recommend either vincristine plus carboplatin or vinblastine monotherapy as first-line treatment. The EAG explained that its clinical advice suggested that vincristine plus carboplatin and vinblastine monotherapy are used interchangeably in clinical practice. So, the EAG felt that vinblastine monotherapy should have been included as a comparator. The clinical experts explained that vincristine plus carboplatin would be first-line systemic treatment for most people. They explained that vinblastine monotherapy would only be considered for first-line treatment in neurofibromatosis type 1 LGG, or for people who cannot tolerate carboplatin. As noted in the CCLG guidelines, vinblastine monotherapy would typically be used as a second-line treatment for people with non-neurofibromatosis type 1 LGG (which includes most people with LGG). The committee concluded that vincristine plus carboplatin was the most appropriate comparator for dabrafenib plus trametinib in LGG.

HGG cohort

3.4 The final NICE scope listed chemotherapy and best supportive care as the comparators for HGG. The company noted that temozolomide is the only chemotherapy that is licensed for children and young people with HGG. Because temozolomide is regularly used as adjuvant treatment with radiotherapy, the company explained that the relevant comparator is dependent on whether people had previously had temozolomide. For

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people with HGG who have not previously had temozolomide, the relevant comparator is temozolomide. For people with HGG who have previously had temozolomide, the relevant comparator is best supportive care. The clinical experts agreed with these comparators. The committee concluded that the most appropriate comparators for HGG were temozolomide (for people who had not previously had temozolomide), and best supportive care (for people who had previously had temozolomide).

Clinical effectiveness

Data sources

3.5 The clinical evidence came from TADPOLE, a multicentre, open-label, phase 2 study done across 20 countries (including the UK). TADPOLE consisted of 2 sub-studies. The LGG cohort was a randomised controlled trial in which people aged 1 to 17 years with BRAF V600E mutationpositive LGG were randomised to have dabrafenib plus trametinib (N=73) or vincristine plus carboplatin (N=37). The HGG cohort was a single-arm prospective cohort study in which people aged 1 to 17 years with BRAF V600E mutation-positive HGG had dabrafenib plus trametinib (N=41). People in both cohorts had treatment until progression, unacceptable toxicity, lack of clinical benefit, start of new treatment or death. The median follow up was 39.0 months in the LGG cohort and 45.2 months in the HGG cohort.

LGG cohort

3.6 The primary outcome measure was overall response rate. In the dabrafenib plus trametinib arm it was 54.8% (95% confidence interval [CI] 42.7% to 66.5%) and in the vincristine plus carboplatin arm it was 16.2% (95% CI 6.2% to 32.0%). Secondary outcomes included progression-free survival and overall survival. Progression was assessed by investigators and by central independent review. Median progression-free survival by investigator assessment was 46.0 months (95% CI 38.6 months to not estimable) in the dabrafenib plus trametinib arm and 30.8 months (95% CI

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7.0 months to not estimable) in the vincristine plus carboplatin arm. Median progression-free survival by independent review was 24.9 months (95% CI 12.9 to 31.6 months) in the dabrafenib plus trametinib arm and 7.2 months (95% CI 2.8 to 11.2 months) in the vincristine plus carboplatin arm. Overall survival data was immature at the end of the study, with 1 death in the vincristine plus carboplatin arm and no deaths in the dabrafenib plus trametinib arm. The committee concluded that the evidence from TADPOLE showed that dabrafenib plus trametinib was more effective than vincristine plus carboplatin for treating LGG.

HGG cohort

3.7 Overall response rate was the primary outcome measure and was 56.1% (95% CI 39.7% to 71.5%) with dabrafenib plus trametinib. Median progression-free survival by investigator assessment was 24.0 months (95% CI 12.5 months to not estimable), and by independent review was 9.0 months (95% CI 5.3 to 20.1 months). Overall survival data was immature at the end of the study, with 17 deaths (41.5%); median overall survival not estimable (95% CI 19.8 months to not estimable). The committee concluded that the evidence from TADPOLE was the best available to show the effectiveness of dabrafenib plus trametinib for treating HGG.

Generalisability of LGG data

3.8 For LGG, dabrafenib plus trametinib is indicated for people who need systemic treatment. The EAG noted that the clinical evidence from the comparative part of TADPOLE was from people who were eligible for firstline systemic treatment. So, there was no comparative evidence or economic analyses of dabrafenib plus trametinib as a treatment for LGG at the second or later lines of treatment. The clinical experts noted that, in current clinical practice, the availability of dabrafenib plus trametinib is restricted by managed access programmes or clinical trials, so it is only used at second and later lines of treatment. They explained that this

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restriction was purely practical and that most clinicians would prefer to use dabrafenib plus trametinib as a first-line treatment for BRAF V600E mutation-positive LGG. Because of this, the committee concluded that the evidence from TADPOLE for first-line use was sufficient to evaluate dabrafenib plus trametinib, because this is the point in the pathway at which it will likely be used for LGG.

Indirect treatment comparison for HGG

3.9 Because the HGG cohort of TADPOLE was single arm, the company did unanchored indirect treatment comparisons (ITCs) to compare dabrafenib plus trametinib with the comparators. For the no prior temozolomide subgroup, the company's systematic literature review identified 2 studies. both of which had used temozolomide. For the prior temozolomide subgroup, the company was unable to identify any studies using best supportive care. The company received clinical advice that after temozolomide has not worked, chemotherapy tends to be ineffective. So, using studies in which people who previously had treatment with temozolomide then had chemotherapy would be a reasonable proxy for best supportive care. The company was able to identify 2 such studies: 1 that used cilengitide and 1 that used bevacizumab. The unanchored ITCs used either matching-adjusted indirect comparison or inverse probability of treatment weighting methods. Matching-adjusted indirect comparison was used when only aggregate data was available for the comparator; inverse probability of treatment weighting was used when patient-level data was available. For both subgroups, the ITCs produced progressionfree survival and overall survival hazard ratios that were less than 1 and had 95% CIs that did not include 1. This implied that dabrafenib plus trametinib statistically significantly improved overall survival and progression-free survival compared with temozolomide and best supportive care. The exact results of the ITCs are considered confidential by the company so cannot be reported here. The EAG agreed with the methods used for the ITCs but cautioned that they are associated with

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uncertainty. The EAG noted the small sample sizes, the limited covariates adjusted and the lack of data on BRAF V600E mutation status in the comparator studies. It noted that for the no prior temozolomide ITC, the comparator studies were approximately 20 years old. It also noted that for the prior temozolomide ITC, the comparator studies used chemotherapy and so were only proxies for best supportive care. The committee acknowledged the uncertainty with using unanchored ITCs to establish comparative efficacy but concluded that they are acceptable for decision-making.

Economic model

Company's modelling approach

- 3.10 The company developed an individual-based state transition model which included analyses for both LGG and HGG cohorts. There were 3 health states common to both analyses: progression-free after first treatment, progressed and death. The LGG part of the model had an additional health state for when people with LGG transform to HGG (secondary HGG). The model simulates the individual histories of a sample of people aged 1 to 17 years with BRAF V600E mutation-positive glioma over a lifetime horizon. The time that people spent in the various health states was based on time-to-event data from TADPOLE and literature sources. The key events, as described in the model, included:
 - 'progression (not because of malignant transformation)
 - malignant transformation (in the LGG analysis)
 - death (glioma-related death, malignant death or non-glioma related)'.

The committee was satisfied with the company's modelling approach.

Progression assessment

3.11 The company and EAG disagreed on whether to use independentassessed or investigator-assessed progression-free survival from

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TADPOLE to model progression. As noted in <u>section 3.6</u> and <u>section 3.7</u>, there were large differences between the results of these assessments of progression-free survival. The company used investigator-assessed progression-free survival in its base case. It received clinical advice that, in clinical practice, investigator-assessed progression-free survival is a more accurate reflection of the timing of progression and of the decision to stop treatment than independent review of progression-free survival. The EAG disagreed and noted that independent review of progression-free survival is more widely used in clinical trials. Its preferred approach would be to use investigator-assessed progression-free survival for treatment discontinuation and independent-assessed progression-free survival for health state occupancy. But, it noted that the company's model was not flexible enough to allow this. So, it suggested a pragmatic approach which used independent-assessed progression-free survival for both. The clinical experts explained that the difference between the investigator and independent estimations of progression-free survival was because independent reviewers only see the tumour scan in isolation. In contrast, investigator assessment accounts for the scan, as well as information directly gained from the person with glioma and their caregiver. The committee agreed with the company and clinical experts that investigator assessment is a more accurate reflection of when progression occurred. It concluded that for this evaluation, investigator-assessed progression-free survival should be used.

Extrapolation

3.12 The company's base case used a piecewise hybrid approach to extrapolate progression-free survival over the entire time horizon. The company justified the piecewise approach on the basis that parametric extrapolations did not fit the observed Kaplan–Meier (KM) curves well. The cut-off point (next to last observed event) was chosen because of the low number of people at risk after 2 years and because it was aligned to the expected treatment duration of dabrafenib plus trametinib in clinical

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practice. The EAG felt that the piecewise approach was not appropriate because the choice of cut-off point was highly arbitrary. It also noted that the company had not adequately justified using the same rate of progression for both arms in the extrapolation phase of LGG, nor using a constant hazard ratio for lifetime in HGG. The EAG's preferred approach was to fit independent parametric curves and extrapolate across the entire time horizon. The committee considered that the company's piecewise approach had limitations. It noted that there was only a small number of people in TADPOLE, and that this meant that there was not much data used to generate the KM curve. The committee thought that this may limit the generalisability of the KM curves from TADPOLE to people with glioma in NHS practice. It also agreed with the EAG that the choice of cutoff point was arbitrary. So, the committee concluded that it preferred the EAG's approach of the 2 methods presented for extrapolating progression-free survival.

Treatment duration

3.13 The marketing authorisations for dabrafenib and trametinib state that treatment should continue until 'disease progression or until the development of unacceptable toxicity'. They also state that 'there are limited data in patients older than 18 years of age with glioma, therefore continued treatment into adulthood should be based on benefits and risks to the individual patient as assessed by the physician'. In the company's base case for LGG, the KM data was used to model progression until week 193 (about 3.7 years). This aligned with clinical advice given to the company that suggested that treatment with dabrafenib plus trametinib would be stopped around 2 to 5 years, in the absence of progression, because of cumulative toxicity. For HGG, clinical advice to the company suggested that clinicians would be more reluctant to stop treatment, given the lack of alternative options and the poor prognosis of HGG. Nevertheless, some people with glioma that has a maintained response to dabrafenib plus trametinib may stop treatment because of cumulative

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toxicity. In the company HGG base case, the model assumes an informal stopping rule of 12.5 years if there is no progression. The EAG noted that the treatment duration of dabrafenib plus trametinib is uncertain and that removing the stopping rule makes it less cost-effective. The clinical experts considered that the likely duration of dabrafenib plus trametinib is difficult to predict. This is because it is likely that there will always be some people whose glioma responds well to treatment and who experience minimal side effects. Furthermore, when stopping dabrafenib plus trametinib, people with glioma may experience high relapse rates and then need retreatment. The committee concluded that treatment duration should reflect the marketing authorisations and a stopping rule should not be included for either LGG or HGG.

Treatment dosing

3.14 The company's base case assumed that dabrafenib plus trametinib was dosed in line with the schedule in TADPOLE, in which both age and weight determined the dose. The marketing authorisation simplifies this by only dosing by weight. So, the company provided a scenario analysis in which the dose was determined by weight. The committee considered that dosing by weight would be how dabrafenib plus trametinib would be used in NHS clinical practice. It concluded that dosing by weight should be used in the model.

Utility values

3.15 The company's systematic review was unable to identify utility values for children and young people with glioma. So, it had to source all values from adults with glioma. The EAG acknowledged the lack of evidence but cautioned that adult utility values may be invalid for children and young people. The clinical experts noted that adult utility values would likely be valid, but they recognised the uncertainty. The committee considered that the adult values likely underestimate the utility decrements that would be seen in children and young people. This is because of the uncaptured

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wider impact on caregivers and families of children and young people with glioma, and the effect of the loss of socialising and educational delays that would be more acutely felt in children and young people. Further, the committee noted the EAG's critique which stated that the use of decrements rather than a multiplicative approach likely reduced the health-related quality of life lost in the model. The committee concluded that the utility values from adults are acceptable to model health-related quality of life but highlighted that the decrements are likely to be an underestimation.

Severity

3.16 The committee considered the severity of the condition (the future health lost by people living with the condition and having standard care in the NHS). The committee may apply a severity modifier (a greater weight to quality-adjusted life-years [QALYs]) if technologies are indicated for conditions with a high degree of severity. The company and EAG provided absolute and proportional QALY shortfall estimates in line with <u>NICE's</u> <u>manual on health technology evaluations</u>. Both the company and the EAG agreed that the QALYs in the LGG population should have a higher weighting (1.2 multiplier), and that QALYs in the HGG population should have the highest weighting (1.7 multiplier). The committee concluded that severity weights of 1.2 (LGG) and 1.7 (HGG) applied to the QALYs were appropriate for this evaluation.

Cost-effectiveness estimates

Acceptable ICER

3.17 <u>NICE's manual on health technology evaluations</u> notes that, above a most plausible incremental cost-effectiveness ratio (ICER) of £20,000 per QALY gained, judgements about the acceptability of a technology as an effective use of NHS resources will take into account the degree of certainty around the ICER. The committee will be more cautious about recommending a technology if it is less certain about the ICERs

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presented. But it will also take into account other aspects including uncaptured health benefits. The committee noted that there were several sources of uncertainty, specifically that:

- the comparative efficacy of dabrafenib plus trametinib in HGG was based on indirect comparison (see <u>section 3.9</u>)
- the progression-free survival extrapolations were uncertain and based on KM data from a small number of people (see <u>section 3.12</u>)
- the likely duration of treatment with dabrafenib plus trametinib that would be used in NHS clinical practice is unclear (see <u>section 3.13</u>)
- the utility decrements used in the model were sourced from adults (see section 3.15).

But, the committee also recalled the statements from the clinical and patient experts that children and young people with BRAF V600E mutation-positive glioma would highly value a new treatment option. It also recalled that they would particularly welcome a treatment that would allow them to live a less restricted life with fewer visits to the hospital. It also noted that because of the rarity of BRAF V600E mutation-positive glioma, the decision risk to the NHS was low. So, the committee concluded that an acceptable ICER would be around £30,000 per QALY.

Company and EAG cost-effectiveness estimates

3.18 <u>NICE's manual on health technology evaluations</u> states that the committee's preferred cost-effectiveness estimates should be derived from a probabilistic analysis when possible unless the model is linear. In this evaluation, the EAG noted that the model was linear and so deterministic analyses were acceptable for decision-making. In the LGG analysis, because of confidential commercial arrangements for other treatments in the model, the exact cost-effectiveness estimates are confidential and cannot be reported here. Both the company's and EAG's

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base case ICERs were within the range that NICE considers to be a costeffective use of NHS resources, but the EAG's was lower. In the HGG analysis, the company's base case deterministic ICER with the severity weighting applied, was £28,624 in the no prior temozolomide subgroup, and £29,072 in the prior temozolomide subgroup. The EAG's base case deterministic ICER, with the severity weighting applied, was £27,500 in the no prior temozolomide subgroup, and £21,512 in the prior temozolomide subgroup.

Committee's preferred assumptions

- 3.19 The committee's preferred assumptions included:
 - using investigator-assessed progression-free survival from the TADPOLE study (see <u>section 3.11</u>)
 - extrapolating progression-free survival by fitting independent curves to the dabrafenib plus trametinib and comparator KM data and extrapolating over the entire time horizon (see <u>section 3.12</u>)
 - not including a stopping rule for modelling treatment duration for either LGG or HGG (see <u>section 3.13</u>)
 - using dosing of dabrafenib and trametinib based on weight (see <u>section</u> <u>3.14</u>)
 - applying a severity weighting of 1.2 to LGG QALYs and 1.7 to HGG QALYs (see <u>section 3.16</u>).

Other factors

Equality

3.20 During scoping, consultees noted that the population in the marketing authorisation is restricted to 'paediatric patients aged 1 year and older' and that this contributes to inequality based on age. Age is a protected characteristic under the Equality Act 2010. Because the committee is only able to make recommendations within the marketing authorisation, it

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concluded that this restriction did not represent an equality issue in its evaluation of clinical and cost effectiveness.

Innovation

3.21 The committee considered whether dabrafenib plus trametinib was innovative. The clinical experts explained that because dabrafenib plus trametinib is a targeted treatment for BRAF V600E mutation-positive glioma, it represents a step-change in current treatment. The patient experts noted that because dabrafenib plus trametinib is an oral treatment, it is more convenient, and that people with glioma and their caregivers would value having fewer hospital visits for intravenous treatment. They noted that dabrafenib plus trametinib allowed children and young people to spend more time with family and take part in more recreational activities. The committee noted the benefits associated with better school attendance and increased socialising that would result from fewer visits to the hospital which would not be captured in adult utility values (see section 3.15). For these reasons, the committee concluded that there were uncaptured benefits in the QALY calculations for which it would account for in its decision making.

Conclusion

- 3.22 The committee considered that it had not been presented with an ICER that reflected its preferred assumptions for LGG or HGG. But, the committee recalled:
 - the range of ICERs presented
 - the unmet need for treatment of the condition
 - the potential value of dabrafenib plus trametinib to people with BRAF V600E mutation-positive glioma, their caregivers and families, and clinicians
 - the benefits uncaptured in the QALY calculations
 - the low decision risk

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• NICE's commitment to take a proportionate approach to appraisals.

When accounting for these factors, the committee was satisfied that its preferred assumptions would result in an ICER within the range that it considered a cost-effective use of NHS resources. So, dabrafenib plus trametinib is recommended.

4 Implementation

- 4.1 Section 7 of the <u>National Institute for Health and Care Excellence</u> (Constitution and Functions) and the Health and Social Care Information <u>Centre (Functions) Regulations 2013</u> requires integrated care boards, NHS England and, with respect to their public health functions, local authorities to comply with the recommendations in this evaluation within 3 months of its date of publication.
- 4.2 Chapter 2 of <u>Appraisal and funding of cancer drugs from July 2016</u> (including the new Cancer Drugs Fund) – A new deal for patients, <u>taxpayers and industry</u> 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 cost comparison evaluation), at which point funding will switch to routine commissioning budgets. The <u>NHS England Cancer Drugs Fund list</u> 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

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technology appraisal guidance 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 draft guidance.

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 glioma and the doctor responsible for their care thinks that dabrafenib plus trametinib is the right treatment, it should be available for use, in line with NICE's recommendations.

5 Evaluation committee members and NICE project team

Evaluation committee members

The 4 technology appraisal committees are standing advisory committees of NICE. This topic was considered by <u>committee A</u>.

Committee members are asked to declare any interests in the technologies being evaluated. If it is considered there is a conflict of interest, the member is excluded from participating further in that evaluation.

The <u>minutes of each evaluation committee meeting</u>, which include the names of the members who attended and their declarations of interests, are posted on the NICE website.

Chair

Radha Todd

Chair, technology appraisal committee A

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NICE project team

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

Tom Palmer Technical lead

Sally Doss Technical adviser

Tom Feist Project manager

ISBN: [to be added at publication]

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