



Regorafenib for previously treated unresectable hepatocellular carcinoma: A Single Technology Appraisal

Addendum: ERG exploratory analyses based on 2016 and 2017 data cut-offs

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This addendum presents the ERG's exploratory analyses using the original RESORCE data cut-off (29th February 2016 cut-off, Table 1) and using the later data-cut off (23rd January 2017, see

Table 2). Additional sensitivity analyses using the later 23rd January 2017 data cut-off are presented in Table 3. As shown in the results, the use of the newer data cut-off reduces the ICER for regorafenib versus BSC.

Table 1: ERG exploratory analyses using 29th February 2016 data-cut-off

| Option | QALYs | Costs | Inc. QALYs | Inc. costs | ICER (per QALY gained) |
|---|-------|-------|------------|------------|------------------------|
| <i>Exploratory analysis 1: Correction of unequivocal model errors and use of alternative unit costs</i> | | | | | |
| Regorafenib | 1.048 | | 0.368 | £12,659 | £34,406 |
| BSC | 0.680 | | - | - | - |
| <i>Exploratory analysis 2: Inclusion of more appropriate general ward bed day cost*</i> | | | | | |
| Regorafenib | 1.048 | | 0.368 | £12,647 | £34,376 |
| BSC | 0.680 | | - | - | - |
| <i>Exploratory analysis 3: Use of full pack dosing*</i> | | | | | |
| Regorafenib | 1.048 | | 0.368 | £15,508 | £42,151 |
| BSC | 0.680 | | - | - | - |
| <i>Exploratory analysis 4: Removal of half-cycle correction for drug acquisition costs*</i> | | | | | |
| Regorafenib | 1.048 | | 0.368 | £13,332 | £36,235 |
| BSC | 0.680 | | - | - | - |
| <i>Exploratory analysis 5: Use of combined 2007 and 2015 survey costs*</i> | | | | | |
| Regorafenib | 1.048 | | 0.368 | £20,297 | £55,166 |
| BSC | 0.680 | | - | - | - |
| <i>Exploratory analysis 6: Use of independent Weibull functions to model OS*</i> | | | | | |
| Regorafenib | 0.896 | | 0.265 | £10,242 | £38,683 |
| BSC | 0.632 | | - | - | - |
| <i>Exploratory analysis 7: Use of a fully extrapolated log logistic time to discontinuation curve (patients on treatment at 29th February 2016 censored, with full pack dosing)*</i> | | | | | |
| Regorafenib | 1.048 | | 0.368 | £21,751 | £59,120 |
| BSC | 0.680 | | - | - | - |
| <i>Exploratory analysis 8: ERG's preferred base case (including all individual amendments)*</i> | | | | | |
| Regorafenib | 0.896 | | 0.265 | £21,468 | £81,081 |
| BSC | 0.632 | | - | - | - |

Table 2: ERG exploratory analyses using 23rd January 2017 data-cut-off

| Option | QALYs | Costs | Inc. QALYs | Inc. costs | ICER (per QALY gained) |
|---|--------------|--------------|-------------------|-------------------|-------------------------------|
| <i>Exploratory analysis 1: Correction of unequivocal model errors and use of alternative unit costs</i> | | | | | |
| Regorafenib | 1.072 | | 0.405 | £13,637 | £33,703 |
| BSC | 0.668 | | - | - | - |
| <i>Exploratory analysis 2: Inclusion of more appropriate general ward bed day cost*</i> | | | | | |
| Regorafenib | 1.072 | | 0.405 | £13,536 | £33,456 |
| BSC | 0.668 | | - | - | - |
| <i>Exploratory analysis 3: Use of full pack dosing*</i> | | | | | |
| Regorafenib | 1.072 | | 0.405 | £16,594 | £41,012 |
| BSC | 0.668 | | - | - | - |
| <i>Exploratory analysis 4: Removal of half-cycle correction for drug acquisition costs*</i> | | | | | |
| Regorafenib | 1.072 | | 0.405 | £14,309 | £35,365 |
| BSC | 0.668 | | - | - | - |
| <i>Exploratory analysis 5: Use of combined 2007 and 2015 survey costs*</i> | | | | | |
| Regorafenib | 1.072 | | 0.405 | £22,099 | £54,619 |
| BSC | 0.668 | | - | - | - |
| <i>Exploratory analysis 6: Use of independent Weibull functions to model OS*</i> | | | | | |
| Regorafenib | 0.967 | | 0.319 | £11,553 | £36,241 |
| BSC | 0.648 | | - | - | - |
| <i>Exploratory analysis 7: Use of a fully extrapolated log logistic time to discontinuation curve (patients on treatment at 29th February 2016 censored, with full pack dosing)*</i> | | | | | |
| Regorafenib | 1.072 | | 0.405 | £22,305 | £55,128 |
| BSC | 0.668 | | - | - | - |
| <i>Exploratory analysis 8: ERG's preferred base case (including all individual amendments)*</i> | | | | | |
| Regorafenib | 0.967 | | 0.319 | £23,768 | £74,559 |
| BSC | 0.648 | | - | - | - |

Table 3: New sensitivity analyses using 23rd January 2017 data-cut-off

| Scenario | Inc. QALYs | Inc. costs | ICER (regorafenib versus BSC) |
|---|-------------------|-------------------|--------------------------------------|
| ERG base case | 0.319 | £23,768 | £74,559 |
| Alternative OS functions | | | |
| OS - exponential | 0.348 | £23,836 | £68,462 |
| OS – log normal | 0.410 | £28,851 | £70,409 |
| OS – log logistic | 0.412 | £28,987 | £70,424 |
| OS – Gompertz | 0.343 | £23,296 | £67,835 |
| OS – generalised gamma | 0.408 | £28,764 | £70,551 |
| Alternative time to treatment discontinuation functions | | | |
| TTTD - exponential | 0.319 | £21,461 | £67,320 |
| TTTD – Weibull | 0.319 | £22,832 | £71,622 |
| TTTD – log normal | 0.319 | £23,977 | £75,214 |
| TTTD – Gompertz | 0.319 | £24,192 | £75,888 |
| Alternative utility values | | | |
| Utilities from SHARP trial | 0.281 | £23,768 | £84,597 |
| Disutility due to progression doubled (state utility=0.715) | 0.311 | £23,768 | £76,441 |
| Disutility due to progression tripled (state utility=0.667) | 0.303 | £23,768 | £78,422 |
| Alternative interpretation of company's resource use survey | | | |
| Number of hospitalisations per month estimated per month assumed to apply to the entire population. | 0.319 | £24,481 | £76,793 |
| Inclusion of dose reductions | | | |
| Indefinite dose reduction to 120mg/day | ■ | ■ | ■ |