

## Appendix H – Economic evidence tables

| Study, Population, Country and Quality  | Data Sources   | Other Comments  | Incremental (EVAR vs. OSR / no repair) |                |          | Conclusions  | Uncertainty   |
|---|--|---|--|----------------|----------|--|---|
|   |  |   | Cost (£)                               | Effect (QALYs) | ICER (£) |  |   |
| <p><b>Michaels et al. (2005)</b><br/>Decision tree model comparing EVAR with OSR (and EVAR with no repair). UK.</p> | <p><u>Effects:</u> EVAR-1 and DREAM studies for operative outcomes. NICE review of non-RCTs for other EVAR outcomes.</p> <p><u>Costs:</u> Intervention, monitoring and reintervention. Tariff costs for primary procedure plus £4500 for EVAR. Other resource use from EUROSTAR registry and assumptions.</p> <p><u>Utilities:</u> Short term recovery decrements (NR), followed by general age-related utility after successful repair.</p> | <p>Cohort: male, 70 years old, 5.5cm AAA.</p> <p>10-year time horizon. 3.5% discount rates. Price year 2003-04.</p> <p>No long-term CV events.</p> <p>General population life expectancy applied after successful repair.</p> | <p><u>EVAR vs. OSR</u><br/>11,449</p>  | 0.10           | 110,000  | <p>'The results of this analysis suggested that, in patients in whom conventional open repair would be an alternative, EVAR provided a slight additional benefit, but at a cost that would not normally be considered appropriate for funding by the NHS.'</p> | <p>EVAR ICER &lt;£20,000 in ~0% of 1000 PSA model runs, compared with OSR.</p> <p>Base case result robust to scenario analyses (e.g. assuming £0 EVAR device cost: ICER &gt;£50,000).</p> |
| <p><b>Partially applicable</b><br/>a</p>  |  |   |  |                |          |  |   |
| <p><b>Potentially serious limitations</b> b,c,d,e</p>   |  |   |  |                |          |  |   |

Key: EVAR, endovascular aneurysm repair; ICER, incremental cost-effectiveness ratio; NR, not reported; OSR, open surgical repair; PSA, probabilistic sensitivity analysis; QALY, quality-adjusted life year.

- a. Only considers infrarenal aneurysms.
- b. Relative effects only available for operative outcomes for EVAR vs. OSR comparison; no randomised data used for 'unfit for OSR' population.
- c. Successful repair effectively considered a 'cure' as patients return to general population life expectancy (long-term data not available at the time of analysis).
- d. Reintervention and complications (endoleak) only modelled for EVAR, and no long-term complications modelled.
- e. 10-year time horizon (15 in scenario analysis); shorter than lifetime, and current long-term EVAR-1 data suggest long-term survival differences.

| Study, Population, Country and Quality   | Data Sources   | Other Comments  | Incremental (EVAR vs. OSR) |                          |                | Conclusions  | Uncertainty  |
|--|--|---|----------------------------|--------------------------|----------------|--|--|
|  |  |   | Cost (£) (95% CI)          | Effect (QALYs) (95% CI)  | ICER (£)       |  |  |
| <p><b>Epstein et al. (2008)</b><br/>Markov model comparing EVAR with OSR based on EVAR-1 patients and data. UK.</p> <p><b>Partially applicable</b> <sup>a</sup></p> <p><b>Potentially serious limitations</b> <sup>b,c,d</sup></p> | <p><u>Effects:</u> EVAR-1 study.</p> <p><u>Costs:</u> EVAR-1 study, NHS reference costs and UK literature.</p> <p><u>Utilities:</u> UK population norms (Kind et al. 1999), 1-month surgery morbidity (EVAR-1), cardiovascular conditions (UK literature).</p> | <p>2-year convergence of EVAR and OSR overall survival, despite 4-year aneurysm-related survival benefit for EVAR. 'Other cause' EVAR mortality catch-up factor applied in the model.</p> <p>Aneurysm-related readmissions modelled. Cardiovascular conditions were MI and stroke.</p> <p>Lifetime horizon, 3.5% discount rate applied to all outcomes.</p> | 3,758<br>(2,439; 5,183)    | -0.02<br>(-0.189; 0.165) | EVAR dominated | <p>'EVAR is unlikely to be cost-effective for all patients within collectively funded healthcare systems.'</p> <p>'EVAR may be cost-effective in a subpopulation of elderly patients fit for open surgery ... if patients maintain this early survival advantage over open surgery.'</p> | <p>EVAR ICER 1.2% likely to be ≤£20,000 per QALY gained.</p> <p>Various scenario analyses. Probability was 14.7% if OSR perioperative mortality was 8% (from 5%); and was 26.2% if the patient was aged 82 (from 74) and differences in cardiovascular event rates were omitted.</p> |

Key: CI, confidence interval; EVAR, endovascular aneurysm repair; ICER, incremental cost-effectiveness ratio; OSR, open surgical repair QALY, quality-adjusted life year; VGNW, Vascular Governance North West; yo, years old.

- a. Only considers infrarenal aneurysms.
- b. Informed by early results from a single study.
- c. Unclear whether difference in aneurysm-related mortality over 4 years is extrapolated to lifetime.
- d. Potential conflict of interest.

| Study, Population, Country and Quality  | Data Sources  | Other Comments  | Incremental (EVAR vs. OSR) |                |          | Conclusions  | Uncertainty   |
|---|---|---|----------------------------|----------------|----------|--|---|
|   |   |   | Cost (£)                   | Effect (QALYs) | ICER (£) |  |   |
| <p><b>Chambers et al. (2009)</b><br/>Markov model comparing EVAR with OSR. UK.</p> <p><b>Partially applicable</b><br/><sup>a</sup></p> <p><b>Potentially serious limitations</b> <sup>b,c,d</sup></p> | <p><u>Effects:</u> Baseline risk equations estimated using IPD from the EUROSTAR study. Relative effects from systematic review (EVAR-1 and DREAM).</p> <p><u>Costs:</u> Intervention, monitoring and readmission. Resource use from EVAR-1. Costs from EVAR-1 and UK sources.</p> <p><u>Utilities:</u> UK population norms (Kind et al. 1999), surgery-related decrements for 6 months (EVAR-1).</p> | <p>Lifetime horizon, 3.5% discount rates, Markov model. Price year 2007.</p> <p>Risk equations constructed to predict operative mortality, post-operative mortality, and readmission. Readmissions are AAA-related only. No long-term CV events.</p> <p>Non-AAA mortality converges after ~3 years. AAA-related mortality benefit of EVAR maintained. Rupture fatality rate assumed 100%.</p> | 2,002                      | 0.041          | 48,990   | <p>'The base-case decision model found that EVAR is not cost-effective on average for patients who are fit for open surgery</p> <p>'If patients can be classified into good, average and poor operative risk, then for patients of most ages and aneurysm sizes, EVAR is cost-effective compared with open repair in patients of poor risk but not cost-effective in patients of good risk.'</p> | <p>EVAR ICER 26.1% likely to be ≤£20,000 per QALY gained. ICER is &lt;£30,000 in patients with subjectively poor operative fitness. ICER &lt;£20,000 where (1) EVAR sustained an overall survival benefit over OSR for the patient's lifetime and (2) unit cost of EVAR equal to OSR, follow-up costs lower and reintervention rates lower.</p> <p>ICER £21-22,000 if EVAR operative mortality odds ratio improved (from 0.35 to 0.25), and if overall mortality rates converge at 8 years (vs. 3 years).</p> |

Key: CI, confidence interval; EVAR, endovascular aneurysm repair; ICER, incremental cost-effectiveness ratio; IPD, individual patient data; OSR, open surgical repair; QALY, quality-adjusted life year.

- a. Only considers infrarenal aneurysms.
- b. Relative effects largely drawn from a single study (EVAR-1).
- c. Impact of long-term non-aneurysm complications not captured by model.
- d. Assumption of maintained AAA-related mortality difference not supported by 15-year EVAR-1 study data.

| Study, Population, Country and Quality  | Data Sources  | Other Comments   | Incremental (EVAR vs. OSR / no repair) |                |                | Conclusions   | Uncertainty  |
|---|---|--|--|----------------|----------------|---|--|
|   |   |  | Cost (£)                               | Effect (QALYs) | ICER (£)       |   |  |
| <b>Brown et al. (2012)</b><br>Markov model comparing EVAR with OSR. Trial analysis comparing EVAR with no repair. UK. | <u>Effects:</u> EVAR-1 and EVAR-2 studies, including ITT analyses.<br><u>Costs:</u> Intervention, monitoring and readmission. Resource use from EVAR trials. Costs from trials and UK sources. In EVAR-2 analysis, costs not extrapolated beyond observed 8-year data.<br><u>Utilities:</u> EVAR-1 analysis: surgery-related decrements for 3 months (EVAR-1 analysis). EVAR-2 analysis: EQ-5D data from trial. | EVAR-1 analysis: Lifetime horizon. EVAR-2 analysis: 8-year analysis and lifetime analysis.<br><br>3.5% discount rates. Price year 2008-09.<br><br>EVAR-1 model: Follow-up divided into first 6 months, 6 months to 4 years, 4 to 8 years, and 8 years onwards. AAA mortality converges after 8 years. Ongoing non-AAA mortality SMR of 1.1 vs. general population (based on EVAR-1 and UKSAT).<br><br>EVAR-2 analysis: 2 analyses presented, 1 ITT (by randomised group) and 1 per protocol (excludes subjects who crossed over from 'no surgery' to intervention).<br><br>No long-term CV events. | <u>EVAR-1</u><br>3,521                 | -0.042         | EVAR dominated | <u>EVAR-1</u><br>'For patients with large AAA, who are deemed anatomically suitable for EVAR and anaesthetically fit for open repair, [EVAR] is a more costly treatment option [than OSR] and unlikely to be cost-effective in all patients.'<br><br><u>EVAR-2</u><br>'For patients deemed anatomically suitable for EVAR but too unfit to for open repair, EVAR offers a long-term benefit in aneurysm mortality ... no benefits in quality of life and high rates of adverse events, complications and reinterventions after EVAR contribute to poor cost-effectiveness.' | <u>EVAR-1</u><br>EVAR ICER 1% likely to be ≤£20,000 per QALY gained compared with OSR. PSA mean costs: £3,519 (95% CI: 1,919 to 5,053). PSA mean QALYs: -0.032 (-0.117 to 0.096).<br>Robust to univariate sensitivity analysis based on alternative clinical data (OVER) and modelling assumptions (Epstein 2008, NICE 2009).<br><br><u>EVAR-2</u><br>0% and 3% of 1000 bootstrapped ICERs were ≤£20,000 (ITT analysis). Mean ICER of lifetime 'per protocol' analysis was £17,805 (61% ≤£20,000). |
|   |   |  | <u>EVAR-2 8-years</u><br>10,214        | 0.037          | 264,900        |   |  |
|   |   |  | <u>Lifetime</u><br>10,214              | 0.350          | 30,274         |   |  |
| <b>Partially applicable</b> <sup>a</sup>  |   |  |  |                |                |   |  |
| <b>Potentially serious limitations</b> <sup>b,c,d</sup>   |   |  |  |                |                |   |  |

Key: CI, confidence interval; EVAR, endovascular aneurysm repair; ICER, incremental cost-effectiveness ratio; ITT, intention to treat; OSR, open surgical repair; PSA, probabilistic sensitivity analysis; QALY, quality-adjusted life year; SMR, standardised mortality ratio; UKSAT, UK Small Aneurysm Trial.

a. Only considers infrarenal aneurysms.

b. Relative effects largely drawn from a single study for each analysis (EVAR-1 and EVAR-2), though these are the only studies to provide ITT data.

c. Impact of long-term non-aneurysm complications not captured by model.

d. Long-term costs not included in the EVAR-2 lifetime extrapolation.

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|--|--|--|--|-------------------------|----------------|--|-------------|
|  |  |  | Cost (£) (95% CI)                        | Effect (QALYs) (95% CI) | ICER (£)       |  |             |
| <b>Epstein et al. (2014)</b><br>Markov model comparing EVAR with OSR based on 4 RCTs. UK.<br><br><b>Partially applicable</b> <sup>a</sup><br><br><b>Potentially serious limitations</b> <sup>b,c</sup> | <u>Effects:</u> EVAR-1, ACE, DREAM and OVER studies.<br><u>Costs:</u> EVAR-1 (UK), ACE (France), DREAM (Netherlands) and OVER (US).<br>Converted to 2009 UK pounds using purchasing power parities.<br><u>Utilities:</u> 3-month surgery morbidity (EVAR-1). | Model based on Epstein et al. (2008) EVAR-1 model. EVAR-1 8-year data used. Cardiovascular complications not modelled.<br><br>4 individual models, no synthesis of RCT data. Each analysis applies the relative survival (including convergence of curves), reintervention data and resource use from the relevant RCT.<br><br>Lifetime horizon, 3.5% discount rate applied to all outcomes. | <u>EVAR-1</u><br>4,014<br>(2,167; 5,942) | -0.02<br>(-0.19, 0.05)  | EVAR dominated | 'This economic analysis does not find that EVAR is cost-effective compared with open repair over the long term based on the EVAR-1, DREAM or ACE trials. EVAR does appear to be cost-effective over the long term based on the OVER trial.'<br><br>EVAR ICER 0% likely to be <£20,000 in the base case EVAR-1, ACE and DREAM analyses, rising to 3% in a favourable scenario.<br><br>EVAR ICER 91% likely to be <£20,000 in the base case OVER analysis, rising to 99% in a favourable scenario. |             |
|  |  |  | <u>ACE</u><br>2,086<br>(1,526; 2,869)    | -0.01<br>(-0.07, 0)     | EVAR dominated |  |             |
|  |  |  | <u>DREAM</u><br>3,181<br>(1,557; 4,986)  | 0<br>(-0.07, 0.05)      | 2,845,315      |  |             |
|  |  |  | <u>OVER</u><br>-1,852<br>(-5,581; 2,097) | 0.05<br>(-0.06, 0.13)   | Dominant       |  |             |

Key: CI, confidence interval; EVAR, endovascular aneurysm repair; ICER, incremental cost-effectiveness ratio; OSR, open surgical repair QALY, quality-adjusted life year; RCT, randomised controlled trial.

a. Only considers infrarenal aneurysms.

b. Each analysis informed by a single study; no synthesis of data.

c. EVAR-1 analysis is very similar to previous models (Epstein et al. 2008; Chambers et al. 2009; Brown et al. 2012); other analyses use non-UK resource use data.