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Cost-effectiveness of home non-invasive ventilation in patients with persistent hypercapnia after an acute exacerbation of COPD in the UK

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ABSTRACT

Home non-invasive mechanical ventilation (HMV) with home oxygen therapy (HOT) in patients with persistent hypercapnia following an acute exacerbation of chronic obstructive pulmonary disease delays hospital readmission. The economic impact of this treatment is unknown. We evaluated the cost-effectiveness of HMV in the UK healthcare system using data from a previously published efficacy trial. Quality-adjusted life-years (QALYs) were computed from EQ-5D-5L. Accounting for all direct patient costs HOT-HMV was £512 (95%CI £36 to £990) more expensive per patient per year than HOT-alone. This small increase in cost was accompanied by increased quality of life leading to an incremental cost-effectiveness ratio of £10 259 per QALY. HOT-HMV was cost-effective in this clinical population. Trial registration number: NCT00990132.

INTRODUCTION

Chronic obstructive pulmonary disease (COPD) remains a common cause of hospital admission, with patients with persistent hypercapnic respiratory failure having worse outcomes.¹ A few studies have evaluated the cost of home non-invasive mechanical ventilation (HMV) with home oxygen therapy (HOT) for patients with COPD with persistent hypercapnia following hospitalisation.² HOT-HMV has been shown clinically efficacy in a previous UK randomised clinical trial.³ We hypothesised HOT-HMV would be cost-effective in the UK.

METHODS

A full description of the trial design can be found with the trial results.³ Patients with persistent hypercapnia ($\text{PaCO}_2 \geq 7 \text{ kPa}$) between 2 weeks and 4 weeks after resolution of acidosis following an admission with an acute exacerbation of COPD were recruited. Patients were randomised to HOT-HMV or HOT. In addition to clinical data, healthcare utilisation, exacerbation frequency and quality-of-life data were collected at each follow-up visit (6 weeks then 3, 6 and 12 months). A detailed description of quality-adjusted life-year (QALY) calculations is provided in online supplemental eMethods. Medical resource utilisation was recorded throughout the trial at routine follow-up, which was reported by patients and verified by electronic health records, where possible. The economic analysis was conducted over 12 months, reflecting the clinical trial. Costs were calculated from 2017 tariff data from a National Health Service perspective (online supplemental eTable 1). Cost-effectiveness was a prespecified outcome of the efficacy trial using an intention-to-treat (ITT) approach. Details of the ITT and per-protocol approach are in online supplemental eMethods. Sensitivity analyses used realistic minimum and maximum costs.

RESULTS

A total of 116 patients were included in the base-case analysis: 57 in the intervention group (HOT-HMV) and 59 in the control group (HOT) (online supplemental eFigure e1). Baseline patient and retention data are provided in online supplemental



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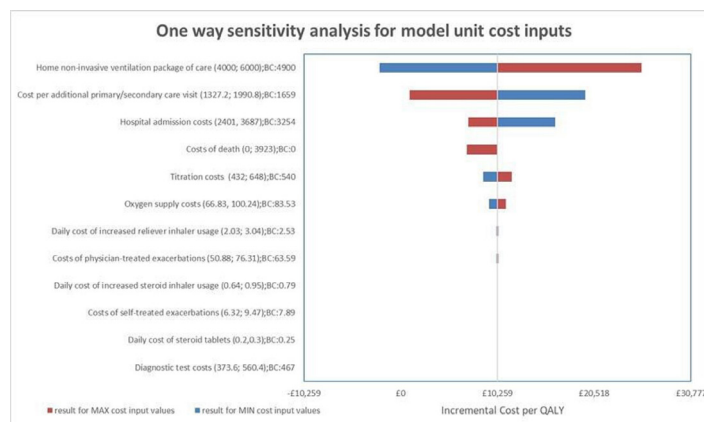


Figure 1 One-way sensitivity analysis results of home non-invasive ventilation with home oxygen therapy versus home oxygen therapy alone in the UK health systems (intention to treat).

Table 1 Cost-effectiveness results for home non-invasive mechanical ventilation (HMV) with home oxygen therapy (HOT) versus HOT alone (intention-to-treat analysis)

Intervention	Total costs (£) (95% CI)	Total QALYs (95% CI)	ICER (Δ cost/ Δ QALYs) (95% CI)
UK analysis			
HOT alone	£16 883 (£13 319 to £20 446)	0.31 (0.23 to 0.39)	Ref
HOT-HMV	£17 395 (£14 309 to £20 482)	0.36 (0.27 to 0.45)	£10 259 (£5438 to £16 449)

ICER, incremental cost-effectiveness ratio; QALYs, quality-adjusted life-years.

eTable 2 and eTable 3. Diary card data were missing for 25 patients (HOT-HMV=8; HOT=17).

Base-case analyses for UK (ITT)

Average total 1-year device costs per patient for the intervention group were £6679 (95% CI £6447 to £6911) compared with £2684 (95% CI £2007 to £3360) in the control group. For all other cost categories, 1-year costs per patient were lower in the intervention group compared with the control group, including average 1-year total primary and secondary care physician visits (£5947 (95% CI £4394 to £7586) vs £8275 (95% CI £6428 to £10 122)); medication costs (£90 (95% CI £52 to £127) vs £104 (95% CI £61 to £146)) and costs for the treatment of exacerbations (£4679 (95% CI £2866 to £6493) vs £5821 (95% CI £4089 to £7552)). Total average annual direct costs per patient were £17395 (95% CI £14309 to £20 482) for the intervention group and £16883 (95% CI £13319 to £20 446) for the control group.

The average number of QALYs was 0.36 (95% CI 0.27 to 0.45) and 0.31 (95% CI 0.23 to 0.39) for the intervention group and control group, respectively. The incremental cost-effectiveness ratio (ICER) was £10 259/QALY (95% CI £5438 to £16 449) (table 1).

One-way sensitivity analyses and bootstrap sensitivity analysis for UK

One-way sensitivity analyses demonstrated parameters with the greatest influence on the ICER: HMV package of care costs for 12 months (ICER range -£2244 to £25 542), cost per additional primary and secondary care physician visit (£944 to £19,574) and hospital admission costs (£7152 to £19574) (figure 1). Bootstrap iterations indicated that at £20 000 and £30 000/QALY, the probability that HOT-HMV is cost-effective versus HOT alone is 56% and 61%, respectively (figure 2A). At £30 000/QALY, the probability that HOT-HMV is more costly and more effective than HOT is 45% (figure 2B). The probability that HOT-HMV is less costly and more effective than HOT alone is 34%.

DISCUSSION

HMV is increasingly used to treat chronic hypercapnic respiratory failure.⁴ A few publications have examined the cost-effectiveness of HMV, with existing economic evaluation largely confined to different HMV modes or setup strategies.⁵⁻⁷ HOT-HMV has previously demonstrated clinical effectiveness, increasing admission-free survival (time to hospital readmission or death) in patients with COPD following a life-threatening exacerbation requiring acute non-invasive ventilation.³ Our study demonstrates that HOT-HMV is cost-effective, with the upper limit of cost per QALY falling below £20 000. This result is consistent with previously conducted analyses using Markov models, which suggest a cost per QALY of £11 318 with a 99% chance of being cost-effective at the £20 000 threshold.^{8,9} The cost per QALY of HOT-HMV falls below that considered acceptable for interventions in the UK¹⁰ and compares favourably with other interventions commonly used in COPD (figure 3).

Limitations

The efficacy trial included a small population with a completion rate (64/116) limited principally by patient death (35/116), which was numerically but not statistically larger in the HOT-alone group. The cost-effectiveness analysis accounts for the lower completion rates as death significantly impacts QALY. Of the patients who withdrew, most (>70%) were in the HOT-alone group. The the most common withdrawal reason was disease progression, which is associated with worse quality of life and so would favour the control arm rather than the intervention.

All centres contributing data to the efficacy trial had established HMV services; therefore, it is possible that less established centres would take longer to set up HMV and would thus increase costs with HOT-HMV. Recent data have demonstrated outpatient setup of HMV in COPD not only is feasible but may be more cost-effective than inpatient titration,⁵ although this was not the case on a recent study of patients with obesity hypoventilation syndrome.¹¹

The trial design allowed patients initially allocated to HOT alone to have HMV if they breached safety criteria after reaching

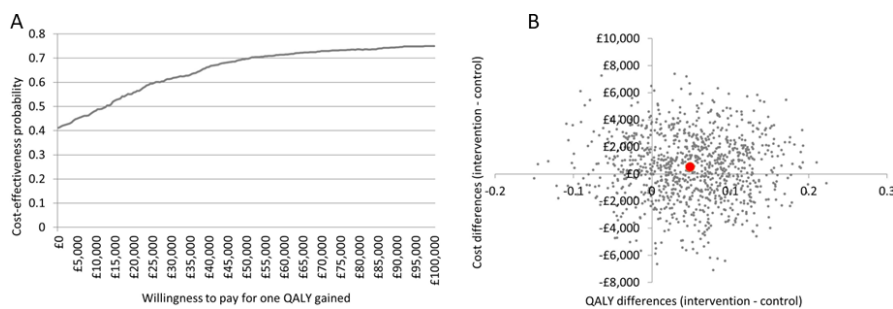


Figure 2 (A) Cost-effectiveness acceptability curve for home non-invasive ventilation with home oxygen therapy versus home oxygen therapy alone in the UK health systems (intention-to-treat analysis); (B) Cost-effectiveness plane for home non-invasive ventilation with home oxygen therapy versus home oxygen therapy alone in the UK health systems (intention-to-treat analysis).

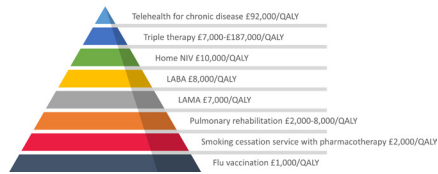


Figure 3 Value pyramid for interventions in the management of COPD. COPD, chronic obstructive pulmonary disease; LABA, long-acting inhaled beta agonist; LAMA, long-acting inhaled muscarinic antagonist; NIV, non-invasive ventilation; QALY, quality-adjusted life-year.

the primary outcome. As expected, the high number of cross-over patients diluted the impact on quality of life between intervention and control arm as these control arm patients were in poorer health than the HOT patients who did not cross-over. Importantly, an additional and modified per-protocol analysis showed increased ICERs compared with ITT (online supplemental eResults).

Finally, the health economic analysis required simplifications. The use of average costs of medical resources does not necessarily reflect actual individual healthcare expenditures but provides typical costs for the patient population. Furthermore, the use of QALYs as an effectiveness measure necessitates breaking the multidimensional construct of quality of life into one value. However, this approach is consistent with other research.¹²

CONCLUSION

HMV with HOT in patients with persistent hypercapnia following an acute life-threatening exacerbation of COPD is likely to be cost-effective in the UK.

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