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Virtual consultations for patients with Obstructive Sleep Apnoea: systematic review and meta-analysis

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Summary of the "take-home" message of the review - 253/256 characters

Virtual consultations are as effective as in-person consultations for follow-up management of adult patients with Obstructive Sleep Apnoea treated with Continuous Positive Airway Pressure. Moreover, this healthcare strategy appears to be cost-effective.

Abstract

Background: The COVID-19 pandemic has accelerated the adoption of virtual care strategies for the management of patients with Obstructive Sleep Apnoea/Hypopnoea Syndrome.

Research question: What is the effectiveness of virtual consultations compared to in-person consultations for the management of Continuous Positive Airway Pressure therapy in adult patients with Obstructive Sleep Apnoea/Hypopnoea Syndrome?

Methods: A systematic review and meta-analysis (PROSPERO; CRD42022297532) based on six electronic databases plus manually selected journals was conducted in January 2022. Two researchers independently selected, quality appraised, and extracted data. The co-primary outcomes were patient-reported sleepiness, assessed by the Epworth Sleepiness Scale, and reported cost-effectiveness.

Findings: 12 studies (n=1823 adults) were included in the review. Seven studies (n=1089) were included in the meta-analysis which showed no difference in the magnitude of improvement in patient-reported sleepiness scores between virtual and in-person consultations (MD [95%CI], -0.39 [-1.38 to 0.60]; p=0.4), though Epworth Sleepiness Scale scores improved in both groups. Virtual care strategies modestly increased Continuous Positive Airway Pressure therapy adherence and were found to be less costly than in-person care strategies in the three Spanish trials that reported cost-effectiveness.

Conclusion: The findings of this review suggest that virtual care delivered by telephone or video consultations is as effective as in-person consultations for improving subjective sleepiness in patients with Obstructive Sleep Apnoea/Hypopnoea Syndrome treated with Continuous Positive Airway Pressure. This clinical management strategy may also improve Continuous Positive Airway Pressure adherence without increasing the costs, supporting its potential as a follow-up management strategy, where patients prefer this approach.

Word count: 248/250 Keywords: Obstructive sleep apnoea (OSA), Sleepiness, virtual consultations, digital health, telemedicine, cost-effectiveness.

Manuscript word count: 2983/3000

Introduction

Obstructive sleep apnoea/hypopnoea syndrome (OSAHS) is estimated to affect nearly 1 billion adults worldwide, with increasing prevalence [1, 2]. OSAHS is associated with debilitating symptoms, reduced neurocognitive performance and quality of life (QoL), increased risk of cardiovascular and metabolic morbidity and occupational accidents [3, 4], and overall represents a major public health concern [5].

Continuous positive airway pressure (CPAP), improves QoL [6]. However, its effectiveness is contingent on optimal adherence [7]. The recently published National Institute for Health and Care Excellence (NICE) guidelines [8] recommend a consultation with the patient within one month of CPAP initiation, and subsequent follow-ups based on patients' needs until optimal outcomes are achieved. This intensive management strategy adds to the strain on respiratory sleep services already coping with increasing demand.

Accelerated by the COVID-19 pandemic [9], digital health interventions such as telemonitoring and virtual consultations have been introduced into sleep medicine to meet this growing demand [10, 11]. Such transition has the potential to revolutionise the way healthcare is delivered, improving accessibility and affordability [12], though there are concerns about exacerbating inequalities, particularly in disadvantaged communities such as minorities and rural populations [13].

In a previous systematic review, we reported on the applicability and feasibility of telemonitoring and virtual consultations in reviewing patients with OSAHS using CPAP. At the time, evidence on clinical and cost-effectiveness of these strategies was scarce [14].

V2.0

Subsequent reviews [15-18] have reported promising findings on the potential of digital interventions to improve adherence to CPAP therapy, but have not evaluated the effectiveness of such strategies on patient-reported outcomes and cost-effectiveness. We, therefore, aimed to systematically review the effectiveness of virtual consultations compared to in-person consultations on patient-reported sleepiness and cost-effectiveness for adults with OSAHS treated with CPAP therapy.

<u>Methods</u>

This systematic review and meta-analysis was registered in the International Prospective Register of Systematic Reviews (PROSPERO; CRD42022297532) and reported in accordance with the Preferred Reporting Items for Systematic Reviews and Meta-Analyses (PRISMA) statement [19].

Search strategy and data sources

A comprehensive search of six electronic databases (MEDLINE, EMBASE, Scopus, Cochrane Library (Central), CINAHL, and medRxiv) was conducted from database inception to 10th January 2022. No limits on publication year or language were imposed. The search strategy, developed in consultation with a medical librarian, used the search terms and keywords: *"Sleep Apnoea, Obstructive", "OSA", "OSAHS", "Sleep-disordered breathing"* AND *"Positive airway pressure", "PAP", "CPAP"* AND *"Virtual consultation", "Remote consultation", "Telemedicine"* (See Supplement, Section 1 for full search strategy).

The search results were de-duplicated in EndNote 20 (Clarivate, Philadelphia, United States). The reference lists of the included studies, other published reviews, and relevant sleep medicine and respiratory journals (e.g. SLEEP, Sleep Medicine, Thorax) were searched manually to identify any additional relevant studies.

Eligibility criteria

The population, intervention, control, and outcomes (PICO) framework for eligibility criteria is summarised in *Table 1*. Studies were included if they were randomised controlled trials (RCTs), quasi-randomised controlled trials (quasi-RCTs), or controlled clinical trials (CCTs) of adult patients with OSAHS using CPAP, comparing virtual consultations to in-person consultations for the follow-up of CPAP therapy.

Study selection and data extraction

We conducted the selection process using Covidence[®] (Veritas Health Innovation, Melbourne, Australia). Two review authors (SA and PM) independently screened titles and abstracts using the predefined inclusion and exclusion criteria. The full-text reports of potentially eligible studies were then assessed independently by the same authors to determine inclusion eligibility. Any discrepancies between reviewers at either stage were resolved by a third review author (JK), or the wider review team (SL, HP, MM, and MP).

Data extraction was performed independently by SA and PM using a customisable form in Covidence[®] software, with comparison and discussion of the findings. The following data were extracted:

- Study design, methodology, follow-up duration, and participants' demographic and baseline data
- Details of intervention, including mode of delivery, intensity, and duration
- Details of comparator
- Relevant findings, specifically those related to our primary and additional outcomes

When needed, corresponding authors were contacted to either seek missing data or clarify unclear methodologies. If the required data were not available in the correct format, the study was excluded from the quantitative meta-analysis.

Risk of bias assessment

The risk of bias (RoB) for each included trial was independently assessed by SA and PM using the Cochrane Collaboration's tool for assessing risk of bias in randomised trials [20]. In the event of a disagreement between the reviewers, a third review author (JK) arbitrated. The assessment of reporting bias through funnel plots was not appropriate in this review due to the small number of studies included in the meta-analyses [21].

Data synthesis

A summary of the included trials is presented, specifically focusing on clinically relevant outcomes including patient-reported sleepiness, cost-effectiveness, and adherence to therapy. Where appropriate data (i.e. mean ± standard deviation) were available for the key outcomes, findings were pooled for a meta-analysis. A random-effects model, with mean differences for continuous data, was performed to pool the results and to calculate 95% confidence intervals and the P values for key outcomes between the virtual consultations group and the control group. The endpoint data after exposure to the intervention were used for the analyses. I² statistic was used to assess the statistical heterogeneity of the included studies; a value greater than 50% was considered an indicator of substantial heterogeneity. Subgroup analyses by the mode of delivery of the virtual intervention and by study follow-up duration were performed. All statistical analyses were conducted using the Cochrane Collaboration's Review Manager Software (RevMan, Version 5.4.1).

<u>Results</u>

Overview of eligible studies

The literature search identified 875 records. After deduplication, 324 studies were retained for initial title and abstract screening. Of these, 63 studies were retrieved for full-text review and assessment for eligibility. 51 studies did not meet the inclusion criteria, and a total of 12 studies [22-33] were included in the review as outlined in the PRISMA flow diagram (**Figure 1**). The commonest reason for exclusion was 'wrong intervention' (n=16). Nine of these studies appeared initially to meet the inclusion criteria, but were excluded because they required an in-person consultation for patients randomised to virtual care; see Supplement, *Section 3* for further details.

Study characteristics

A summary of the included studies is presented in *Table 2*. 11 of the 12 studies were RCTs and one was a CCT, published between 2006 and 2021. These trials had study sample sizes ranging between 45 to 306 participants and follow-up durations spanning 30 days to 6 months. In these trials, multimodal digital health interventions were used to deliver virtual consultations to patients with OSAHS using CPAP. In total, the 12 included studies represented 1823 adult participants with the majority being male (averaging 78%) and overweight, diagnosed with moderate to severe OSAHS (*Table 3*). Only four studies reported ethnicity data [22, 23, 25, 33], and two [22, 32] reported only minimal data on socioeconomic status.

Risk of bias and quality of evidence assessment

A summary of the RoB assessment for the included studies is provided in **Figure 2**. Overall, the assessment showed variation in the RoB among the included studies due to the complex

nature of the intervention. As would be expected, there was a high RoB in all included studies, because of the inability to blind participants and personnel to allocation. Additionally, the RoB arising from allocation concealment was found to be unclear. Imprecision of the findings is unlikely due to the large sample size in the meta-analysis. However, the certainty of evidence was rated down because of the indirectness in the studies. There was variation in how the intervention of interest, virtual consultations, were delivered. Additionally, all the trials recruited the population of interest (patients with OSAHS using CPAP) and investigated the outcomes of interest. Taken together, with reference to the Grading Recommendations, Assessment, Development and Evaluations (GRADE) framework, the evidence to support the use of virtual consultations for improving the primary outcome, ESS scores, in patients with OSAHS is moderate.

Primary outcomes

Patient-reported sleepiness

The change in subjective sleepiness, assessed by ESS, was reported by eight studies [23-25, 28-31, 33]. These studies found an improvement in ESS scores from baseline to follow-up in both virtual and in-person consultation groups, with two trials [29, 30] reporting a significantly greater reduction in ESS scores in the virtual consultation group. No difference was observed in the ESS scores for virtual compared to in-person consultations (MD [95% CI], -0.39 [-1.38 to 0.60]; p= 0.4; moderate-certainty evidence) in the seven studies (n=1089) that had the endpoint data in the correct format for a meta-analysis (**Figure 3**). However, considerable statistical heterogeneity was observed (I²=72%). Subgroup analysis by the mode of delivery of the virtual intervention (telephone alone or video and/or telephone

consultations) and by study follow-up duration (\leq 3 months or 6 months) showed no differences between the study groups (see Supplement, *Section 4.1A and 4.1B*).

Cost-effectiveness

Three trials [24, 27, 28] from Spain investigated the cost-effectiveness of virtual compared to in-person care for patients with OSAHS, from patient and provider perspectives. Isetta *et al.* [24] reported that a telemedicine-based strategy had a slightly lower total cost, with a 69% estimated probability that it would be cheaper than in-person care. When considering only OSA-related costs, the probability increased to 98% with estimated costs of €150.90 and €114.00 respectively for the in-person and virtual care strategies. In another RCT, Turino *et al.* [27] found the total average cost for each randomised participant to be 28% lower in the virtual care group (€123.60) than in the in-person care group (€170.90), with a €47.32 difference between the two study groups. The investigators also reported the Incremental Cost-effectiveness Ratio (ICER) per Quality-Adjusted Life Year which was estimated at €17,358.65 (QALY). The third study conducted by Lugo et al. [28] showed, using Bayesian analysis, that the telemedicine-based strategy was cost-effective; OSA-related costs for the virtual and in-person care groups were €264.96 and €412.03, respectively.

Secondary outcomes

Adherence to CPAP therapy

CPAP therapy usage, assessed objectively, was reported by all included studies (n=12). The average CPAP usage at follow-up assessment ranged from 3.5 to 5.6 hours/night with virtual consultations and 2.1 to 5.6 hours/night with in-person consultations. Nine studies [22-25, 27, 28, 31-33] found no statistically significant difference in the mean hours of CPAP usage between the two care strategies, while the other three studies [26, 29, 30] showed a significantly higher adherence among participants randomised to virtual consultations. The

pooled analysis of 10 RTCs [22-25, 27-31, 33] comprising 1299 participants demonstrated a significant difference in favour of virtual consultations (MD [95% CI], +0.43 [0.06 to 0.80] hours/night; p= 0.02; moderate-certainty evidence); see **Figure 4.** This equates to an average of 26 minutes increase in CPAP usage per night.

Subgroup analysis by the mode of delivery of the virtual intervention (telephone alone or video and/or telephone consultations) and by study follow-up duration (\leq 3 months or 6 months) were performed. The analyses showed attenuation of the effect size in studies with 6 months follow-up compared to studies with \leq 3 months follow-up durations (see Supplement, *Section 4.2A and 4.2B*).

Change in quality of life

10 trials [22-25, 27-31, 33] reported the change in Quality of Life (QoL) from baseline using multiple different generic and disease-specific instruments, including EQ-5D (n=2), The 12-Item Short-Form Health Survey (SF-12) (n=3), the Functional Outcomes of Sleep Questionnaire (FOSQ) (n=4), and a modified FOSQ (n=1). Seven of these trials reported a general improvement in QoL among participants randomised to both virtual and in-person consultations, with no statistically significant differences between the two groups (EQ-5D n=1/2; SF-12 n=2/3; FOSQ n=3/4, M-FOSQ n=1/1). Three trials found a greater improvement of QoL in the virtual consultations group compared to the in-person consultations group (EQ-VAS n=1/2; SF-12 physical component n=1/3; FOSQ n=1/4).

Environmental impact outcomes

None of the trials reported environmental impact outcomes (including travel distances from home to clinic, mode of transport, and carbon-footprint emissions).

Sensitivity analyses

The substantial heterogeneity among the included studies in the meta-analyses prompted post-hoc sensitivity analyses. Nilius et al. [29] met our inclusion criteria, but was thought on clinical grounds to be an outlier, due to its specific patient population (OSA patients with recent ischemic stroke), which may have resulted in unexpected between-groups differences in the study outcomes, specifically ESS scores and CPAP adherence. However, when this study was removed from the meta-analyses, the results remained unchanged: the pooled weighted mean difference for subjective sleepiness was (MD [95% CI], -0.09 [-1.07 to 0.89]; p= 0.86) and for CPAP usage was (MD [95% CI], +0.30 [0.00 to 0.60] hours/night; p= 0.05).

Deviations from protocol

Although the heterogeneity assessment was higher than specified in the protocol (I² >50%), we decided to perform meta-analyses due to the complex nature of the intervention. We have also conducted subgroup and sensitivity analyses.

Discussion

To our knowledge, this is the first systematic review and meta-analysis to evaluate the effectiveness of virtual compared to in-person consultations on both patient-reported sleepiness and also cost-effectiveness. The main findings are that virtual consultations using telephone or videoconferencing are as effective as in-person consultations for improving patient-reported sleepiness (moderate-certainty evidence) in patients with OSAHS treated with CPAP and appear to be cost-effective. Additionally, virtual consultations, modestly increased the average CPAP usage by nearly half an hour per night when compared to in-person consultations. No published studies compared the environmental impact of virtual versus in-person follow-up strategies.

Subjective sleepiness, assessed by the ESS, is an important outcome for the management and monitoring of patients with OSAHS [8]. In this meta-analysis, there was no difference in the magnitude of improvement of ESS scores between virtual and in-person consultations, though ESS decreased in both groups. Importantly, this observation persisted with subgroup analyses by the mode of delivery of the virtual consultation (telephone alone or video and/or telephone consultations) and by study follow-up duration (\leq 3 months or 6 months), suggesting that virtual care delivered in a variety of strategies remained as effective as in-person care.

CPAP adherence was higher in the virtual consultations group than the in-person consultations group, although the 26 minute increase in CPAP usage is slightly lower than the minimal clinically important improvement of 30 minutes per night [34]. This increase in CPAP usage is consistent with earlier meta-analyses which focused on the use of virtual care strategies specifically to investigate the effect on CPAP adherence [15-17]. However, recent post-COVID data has shown a reduction in CPAP adherence with remote CPAP set-up, compared to in-person [11, 35]. While CPAP set-up is just one part of the process of initiation onto therapy, these data highlight the need for robust assessment of new healthcare strategies, before they become mainstream. In the current review, assessment of whether specific characteristics of the virtual consultation could have influenced the adherence to CPAP were performed. Subgroup analysis by follow-up duration showed a trend for the effect size to attenuate in studies with 6 months compared to \leq 3 months follow-up durations, which may support the efficacy of long-term follow-up interventions [36].

Virtual care strategies were less costly than in-person strategies in the three studies that reported cost-effectiveness [24, 27, 28]. Whilst these data need to be interpreted with

caution, it is reassuring that there is no suggestion that virtual follow-up is more costly. The cost savings were mainly driven by fewer in-person follow-up visits, savings on transport, and less loss of productivity. However, the cost-utility analyses were limited to one healthcare context (i.e. Spain) and there was little clarity in how all the calculations of the mean cost per patient were performed. Further global data is likely to emerge as such virtual strategies are continued following the COVID-19 pandemic. Well-designed studies are important to establish whether virtual consultations are a clinically and cost-effective strategy for the management of patients with OSAHS using CPAP, particularly from the patient, societal, and healthcare provider perspectives.

There has been a growing interest in the importance of reporting ethnicity and socioeconomic status data in clinical trials as well as addressing their potential effect on health inequalities [37]. A concerning finding of the current review is that only one-third of the studies reported data on ethnicity and two studies presented minimal data on socioeconomic status. This reduces interpretability and generalisability of their findings, emphasising the need for explicit reporting of trial participants' demographics. Such data facilitates investigations of OSAHS phenotypes, where heterogeneous clinical manifestations exist. This data will also inform discussions around health inequalities as we move towards personalised medicine approaches for OSAHS care [38].

Virtual care strategies have the potential to improve access to healthcare services and address geographical barriers to delivering high-quality care [39, 40]. However, patient-focused concerns exist, specifically about increasing health inequalities ('the digital divide') [41], emphasising the need to ensure that the care is tailored to patients' context, needs and preferences.

One unanticipated finding in this review was that none of the included studies reported data on any of the environmental impact outcomes, despite increased awareness of the climate emergency [42]. This observation signifies a potential gap in evidence related to virtual care strategies for the management of OSAHS. A critical and interesting direction for future research would be to assess the impact of virtual compared to in-person consultations on the environment, reinforcing the need for sustainable delivery of healthcare.

Strengths and limitations

The review was conducted in accordance with current recommendations and guidelines [19, 21]. However, there are several limitations to consider when interpreting the findings. The studies were relatively small in size, with most containing between 100 and 200 participants. They were also varied in terms of the methodologic quality, identified by the RoB assessment. The studies were heterogeneous in type, intensity, and duration of both virtual and in-person care strategies; potentially limiting the interpretation of the results. Care should also be taken when interpreting the subgroup analyses due to the small number of included studies. We suggest future studies should, therefore, be powered to evaluate the different techniques, cost-effectiveness, and the environmental impact of delivering virtual consultations for the management of patients with OSAHS. This is to identify the most efficacious components of these virtual care strategies from patient, clinical, and economic perspectives. Finally, it is important to stress that our analysis did not address the diagnosis of OSAHS, and therefore cannot be used to support a virtual diagnostic pathway.

Conclusion

Our findings suggest that virtual patient care delivered by telephone or video consultations is as effective as in-person consultations for improving subjective sleepiness in patients with

OSAHS treated with CPAP. Moreover, virtual consultations modestly increased CPAP adherence, compared to in-person consultations, and were not associated with reduced cost-effectiveness. Virtual follow-up of patients with OSAHS using CPAP should be available as an alternative care strategy to in-person follow-up, where patients prefer it.

Table 1 PICO	framework for inclusion and exclusion criteria
Population	Adults with a clinical diagnosis of OSAHS, either naïve or established users of CPAP; recruited from any healthcare or community setting. Studies were excluded if they investigated patients with other types of sleep disorders such as Central Sleep Apnoea.
Intervention	Any form of a virtual consultation between a patient and a healthcare provider. This included either synchronous or asynchronous communications made via telephone or videoconferencing, with or without real-time telemonitoring of CPAP. No limitations were imposed regarding the number of consultations, methods of CPAP initiation, or the duration of consultations. Trials that investigated automated interventions, without direct input from a healthcare professional, were excluded.
Comparator	The comparator group were allocated to a clinical in-person consultation, with or without real-time telemonitoring of CPAP therapy.
Outcomes	
	The primary clinical outcome was subjective sleepiness, assessed by the Epworth Sleepiness Scale, and the primary organisational outcome was cost-effectiveness of the intervention.
	Additional outcomes were clinical, patient and/or clinician-reported, and environmental impact outcomes (<i>see Supplement, Section 2 for definitions</i>).
Study design	Studies were included if they were randomised controlled trials (RCTs), quasi-randomised controlled trials (quasi-RCTs), or controlled clinical trials (CCTs)
OSAHS; Obstructive	e Sleep Apnoea/Hypopnoea Syndrome, CPAP; Continuous Positive Airway Pressure

Study (Country)	Study design and overall risk of bias	Number of participants (n=1823)	Population description	Intervention	Intensity and duration	Mode of delivery	Main findings
Taylor et al 2006 [22] (USA)	RCT FU: 30 days High RoB	Total: 114 VC: 56 IP: 58	Adult patients with OSAS who were initiating CPAP therapy	Telemonitoring via the Health Buddy OSAS + Patients with 'high-risk' responses were contacted within 24 h	Patients were contacted as needed to resolve issues	Telephone consultation	ESS was not reported for post- intervention follow-up
Stepnowsky et al 2007 [23] (USA)	RCT FU: 2 months Moderate RoB	Total: 45 VC: 24 IP: 21	Adult patients newly diagnosed with OSA	Telemonitoring via a flow generator data + Objective and subjective patient reports triggered patient contact	Patients were contacted as needed based on a pre- defined clinical pathway	<i>Telephone</i> <i>consultation</i>	No significant differences in ESS scores between the study groups at baseline and post-intervention
Isetta et al 2015 [24] (Spain)	RCT FU: 6 months Moderate RoB	Total: 139 VC: 69 IP: 70	Adult OSA patients requiring CPAP treatment	Telemonitoring via a website developed for this study + input evaluation triggered patient contact	Virtual consultations via skype were scheduled at 1 and 3 months Consultation duration: 38.97 ± 12.04 mins	Video consultation	Improvement in ESS at 6 months, but no significant difference in change from baseline between the study groups The telemedicine-based strategy had a lower total cost compared to standard care
Frasnelli et al 2015 [26] (Switzerland)	CCT FU: 30 days High RoB	Total: 223 VC: 113 IP: 110	Adult patients with sleep apnoea	Telemonitoring via CPAP + A colour-coded algorithm triggered patient contact	Patients were contacted as needed for a duration of ~30 mins	<i>Telephone</i> consultation	ESS was not reported for post- intervention follow-up

Fields et al 2016 [25] (USA)	RCT FU: 3 months Moderate to high RoB	Total: 60 VC: 32 IP: 28	Adult patients with OSA from two community-based outpatient centres	<i>Telemonitoring via APAP</i> + scheduled follow-up contact and if needed	Initial evaluation visit for 40 min with a 10 min (or less) follow up call at week 1. Virtual consultations scheduled at 1 and 3 months for 20 mins each	Initial evaluation via real-time CVT + telephone consultation for follow up	No significant difference in the change of ESS scores from baseline to 3-months follow-up between the study groups
Turino et al 2016 [27] (Spain)	RCT FU: 1 and 3 months Moderate RoB	Total: 100 VC: 52 IP: 48	Adult patients with newly diagnosed OSA requiring treatment with CPAP	Telemonitoring via MyOSA – Oxigen Salud web database + Automatic alarms triggered patient contact	Patients were contacted as needed to resolve issues	<i>Telephone</i> <i>consultation</i>	ESS was not reported for post- intervention follow-up The total average cost per randomised patient was 28% lower in the VC group than in the IP standard care group
Lugo et al 2019 [28] (Spain)	RCT FU: 3 months Moderate RoB	<i>Total: 186</i> <i>VC: 94</i> <i>(32 with</i> <i>CPAP)</i> <i>IP: 92</i> <i>(40 with</i> <i>CPAP)</i>	Adult patients with suspected OSA who were referred to the Sleep Unit	Telemonitoring via CPAP and + input in a custom web application triggered patient contact	Virtual consultations were scheduled at 3, 6, 12 weeks for no more than 15 mins each	Video or Telephone consultation	No significant differences in the ESS scores between the study groups The costs of the VC were cheaper than those for IP standard care, and the Bayesian analysis showed that the VC was cost-effective
Nilius et al 2019 [29] (Germany)	RCT FU: 6 months Moderate to high RoB	Total: 80 VC: 40 IP: 40	Adult OSA patients who had suffered an ischemic stroke within the last 3 months	Telemonitoring + A colour-coded algorithm triggered a more detailed evaluation and patient contact if needed	Patients were contacted as needed for a duration of 5 mins	<i>Telephone</i> <i>consultation</i>	VC group had a significantly lower ESS scores at 6-months follow up

Pepin et al 2019 [30] (France)	RCT FU: 6 month Moderate RoB	Total: 306 VC: 157 IP: 149	Adult patients with severe OSA and high cardiovascular risk	<i>Telemonitoring via CPAP</i> and the multimodal system + Automatic algorithms triggered patient contact	Patients were contacted as needed + regular assessments at day 8 and months 1 and 6	Telephone or teleconsultation	ESS scores significantly improved in both study groups, but the size of improvement was significantly higher in the VC group
Tamisier et al 2020 [31] (France)	RCT FU: 6 months Moderate RoB	Total:206 VC: 102 IP: 104	Newly diagnosed adult patients with OSA and low cardiovascular risk who were referred for CPAP therapy.	Telemonitoring via CPAP and the multimodal system + Automatic algorithms triggered patient contact	Patients were contacted as needed based on an automatic algorithm	Telephone or teleconsultation	ESS scores significantly improved in both study groups, with no significant difference between the groups.
Fietze et al 2021 [33] (Germany)	RCT FU: 6 months Moderate to high RoB	Total: 224 VC: 110 IP: 114	Adult patients with moderate to severe OSA	<i>Telemonitoring via APAP</i> + predefined criteria triggered patient contact	Patients were contacted as needed based on predefined criteria	<i>Telephone</i> consultation	Change from baseline to 6 months in ESS scores was not significantly different between the two groups
Kooij et al 2021 [32] (Netherlands)	RCT FU: 4 weeks 12 weeks 24 weeks Moderate RoB	Total: 140 VC: 70 IP: 70	Adult patients diagnosed with moderate or severe OSA who require CPAP treatment	Telemonitoring + not achieving pre-defined objectives (e.g adherence and residual AHI) triggered patient contact	Patients were contacted as needed + scheduled follow-ups at 1 and 4 weeks	Video and Telephone consultation	ESS was not reported for post- intervention follow-up

RCT; Randomised Controlled Trial, CCT; Controlled Clinical Trial, FU; Follow up duration, RoB; Risk of Bias, ESS, Epworth Sleepiness Scale, VC; Virtual consultations, IP; In-person consultations, OSA; Obstructive Sleep Apnoea, AHI; Apnoea-Hypopnoea Index, CPAP; Continuous Positive Airway Pressure

Table 3 Participant baseline characteristics

Study	Number of participants (n=1823)	Age (years)		Gender	Gender (Male %)		BMI (kg/m2)		AHI (events/hour)	
		Virtual	In-person	Virtual	In-person	Virtual	In-person	Virtual	In-person	
Taylor et al 2006 [22]	Total: 114 VC: 56 IP: 58	45.8 ± 10	44.6 ± 8.5	66%	71%	NR	NR	NR	NR	
Stepnowsky et al 2007 [23]	Total: 45 VC: 24 IP: 21	60 ± 10.8	58 ± 13.7	98%	98%	33.3 ± 4.9	30.5 ± 5.1	44.8 ± 17.9	37.6 ± 14.3	
Isetta et al 2015 [24]	Total: 139 VC: 69 IP: 70	51.0 ± 8.9	47.0 ± 10.9 ¶	85%	87%	32.8 ± 7.3	33.6 ± 8.3	45 (35; 70) [#]	52 (35; 62) [#]	
Frasnelli et al 2015 [26]	Total: 223 VC: 113 IP: 110	55 (47; 63) [#]	55 (45; 61) [#]	76%	78%	NR	NR	37 (23; 62) [#]	40 (20; 69) #	
Fields et al 2016 [25]	Total: 60 VC: 32 IP: 28	46.7 ± 13.1	58.2 ± 14.4	93%	94%	33.2 ± 6.0	32.9 ± 3.8	32.0 ± 33.9	30.1 ± 30.0	
Turino et al 2016 [27]	Total: 100 VC: 52 IP: 48	56 ±13	54 ±12	77%	77%	35 ± 7	35 ± 7	52 ± 25	53 ± 26	
Lugo et al 2019 [28]	Total: 186 VC: 94 (32 with CPAP) IP: 92 (40 with CPAP)	50.39 ± 11.31	50.82 ± 12.15	70%	66%	29.97 ± 6.19	31.50 ± 10.91	24.68 ± 21.01	33.60 ± 28.96	
Nilius et al 2019 [29]	Total: 80 VC: 40 IP: 40	55.4 ± 10.4	58.6 ± 9.3	73%	73%	31.7 ± 5.4	<i>30.1</i> ± 6.6	41.2 ± 19.0	37.6 ± 18.4	

Pepin et al 2019 [30]	Total: 306 VC: 157 IP: 149	60.8 (53.8; 66) [#]	61.8 (54.7; 66.1) [#]	73%	74%	32.4 (29.6; 36.5) [#]	31.4 (28.1; 35) [#]	47 (35; 60.5) [#]	45 (35.4; 61.2) #
Tamisier et al 2020 [31]	Total:206 VC: 102 IP: 104	51.9 (44.8; 58.8) *	49.4 (40; 57.9) [#]	58%	68.3%	30.5 (27.5; 34.9) *	31.3 (26.6; 35.2) [#]	45 (34; 58) [#]	42.3 (33; 57.6) #
Fietze et al 2021 [33]	Total: 224 VC: 110 IP: 114	53.6 ± 11.8	53.1 ± 10.6	83%	79%	32.8 ± 6.4	31.7 ± 5.6	35.3 ± 17.6	37.0 ± 20.3
Kooij et al 2021 [32]	Total: 140 VC: 70 IP: 70	52.3 ± 12.4	54.3 ± 11.9	83%	76%	BMI > 30: n =51 (73%)	BMI > 30 n=46 (66%)	31 (22; 46) [#]	30.5 (20; 42) [#]

Data are presented as \mathbf{n} or $\mathbf{mean} \pm \mathbf{sd}$, unless otherwise stated. VC; Virtual Consultation, IP; In-person consultation, BMI; Body Mass Index AHI; Apnoea-Hypopnoea Index, NR; Not reported. [#]; median (interquartile range), [¶]; Significantly different value from the other group.

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