BMJ Open Long-term use benefits of personal frequency-modulated systems for speech in noise perception in patients with stroke with auditory processing deficits: a non-randomised controlled trial study

Nehzat Koohi,^{1,2} Deborah Vickers,³ Jason Warren,^{2,4} David Werring,^{2,5} Doris-Eva Bamiou^{1,2}

ABSTRACT

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For numbered affiliations see end of article.

Correspondence to Dr Doris-Eva Bamiou; d.bamiou@ucl.ac.uk **Objectives:** Approximately one in five stroke survivors suffer from difficulties with speech reception in noise, despite normal audiometry. These deficits are treatable with personal frequency-modulated systems (FMs). This study aimed to evaluate long-term benefits in speech reception in noise, after daily 10-week use of personal FMs, in non-aphasic patients with stroke with auditory processing deficits.

Design: This was a prospective non-randomised controlled trial study. Patients were allocated to an intervention care group or standard care subjects group according to their willingness to use the intervention or not.

Setting: Tertiary care setting.

Participants: Nine non-aphasic subjects with ischaemic stroke, normal/near-normal audiometry and auditory processing deficits and with reported difficulties understanding speech in background noise were recruited in the subacute stroke stage (3–12 months after stroke).

Interventions: Four patients (intervention care subjects) used the FMs in their daily life over 10 weeks. Five patients (standard care subjects) received standard care.

Primary outcome measures: All subjects were tested at baseline (visit 1) and 10 weeks later (visit 2) on a sentences in noise test with the FMs (aided) and without the FMs (unaided).

Results: Speech reception thresholds showed clinically and statistically significant improvements in intervention but not in standard care subjects at 10 weeks in aided and unaided conditions.

Conclusions: 10-week use of FMs by adult patients with stroke may lead to benefits in unaided speech in noise perception. Our findings may indicate auditory plasticity type changes and require further investigation.

Trial registration number: Pre-results; NCT02889107.

Strengths and limitations of this study

- This is the first study to investigate long-term benefits of frequency-modulated systems use in stroke subjects in order to determine potential value in running a large-scale intervention trial.
- Recruited patients had detailed auditory assessment.
- The study has small numbers and lacks randomisation.
- Extent of lesion effects was not investigated.

INTRODUCTION

Stroke can affect all levels of the auditory pathway¹ and manifests with pure-tone detection deficits on audiometry,^{2 ³} auditory processing deficits^{4 ⁵} and perceptual difficulties in the domains of speech, sound recognition and localisation.⁶ Approximately one in five of stroke survivors report severe difficulties with speech recognition in the presence of background noise, despite the presence of normal audiometry.⁷ These deficits are treatable with personal frequency-modulated (FM) systems, with a robust immediate improvement of 9 dB in signal-to-noise ratio (SNR)when using the FM system.⁷ This is a clinically significant improvement as 6 dB is the cut-off value for the patient to seek clinical intervention.⁸

Personal FM systems are wireless listening devices that pick up the speaker's voice and transmit it to a receiver in the listener's ear, thus reducing the negative effects of noise, distance and reverberation. FM systems are used to improve speech in noise perception in patients with listening difficulties due to disordered auditory processing, such as children with developmental disorders⁹ ¹⁰ and adults with neurological disorders¹¹ ¹² including stroke⁷ with good immediate benefits. Long-term benefits of FM systems have predominantly been investigated on paediatric populations. Children with disordered auditory processing who used FMs 5 months in the classroom showed an improved unaided (ie, with no FM device) speech in quiet performance by 3.8 dB, suggesting the possibility of bottom-up driven auditory neuroplasticity;⁹ however, due to the lack of controls, maturation effects could not be entirely excluded. Dyslexic children who used FMs over 1 year similarly showed improved phonological awareness and reduced variability of subcortical responses to sound.¹⁰

The potential neuroplasticity-type benefits of prolonged FM system use are of particular interest for stroke survivors with acquired auditory processing deficits. The central auditory nervous system maintains the capacity to be altered in response to auditory stimulation or deprivation throughout life.¹³ Language-based rehabilitation leads to plasticity-type benefits for aphasic stroke subjects,¹⁴ and it would be reasonable to expect similar benefits in less impaired (ie, non-aphasic) patients with stroke with speech reception impairments who receive an intervention that promotes better access to speech.

The present study thus aimed to evaluate the potential benefits in speech perception of personal FM systems, when used daily over 10 weeks by non-aphasic patients with stroke with auditory processing deficits, in order to investigate whether plasticity occurs after prolonged use of FM systems.

We hypothesised that improvement in unaided speech in background noise performance, shown on a behavioural task, would occur in patients with stroke who used the FM systems for a period of time, but not in a control group of patients with stroke who did not use these and received standard care.

METHODS

Consent

Written informed consent was obtained from all participants. This study was registered with ClinicalTrials.gov (NCT02889107).

Study design

This study was conducted prospectively with a nonrandomised controlled design. We further aimed to establish how many eligible patients would be willing to use the FMs daily, in order to inform a subsequent trial, the SD of the primary outcome measure, in order to calculate sample size for a subsequent trial and the number of patients willing to use this intervention, follow-up rate and adherence/compliance rate to the intervention.

Participants of our previous study⁷ were asked whether they would be willing to use the FM systems at home for 10 weeks. Four out of nine (44%) agreed to use the FM (subjects 1, 4, 6 and 9—table 1) and formed the intervention subjects group, while five out of nine (56%) who did not wish to use the FM but were willing to come back for a reassessment 10 weeks later formed the standard care group (subjects 2, 3, 5, 7 and 8—table 1).

Participants

All recruited subjects were right-handed adults (age range 24–78 years) with a clinical history of ischaemic stroke verified by brain MRI in the subacute stroke stage (table 1). They had normal/near-normal puretone audiogram (PTA) average thresholds (better than 25 dB hearing level) and reported difficulties on the speech in noise subscale of the Amsterdam Inventory for Auditory Disability (AIAD) (z score >2).⁶ They had abnormal performance (in one ear or both) in the speech in babble¹⁵ and in at least one more non-speech auditory processing test.¹⁶ ¹⁷ No subject had severe aphasia (cut-off of 93.8 on the Western Aphasia Battery (WAB) test¹⁸), significant psychiatric illnesses, other neurological disorders except stroke or severe concurrent medical illnesses.

Baseline assessments

All participants had a brain MRI within 48 hours after the stroke. They had audiological and other baseline assessments over a single test session after recruitment, at 3–12 months after stroke (subacute stage), that included:

Table 1 Lesion description, age, sex, PTA (average in dB HL at 500, 1000, 2000 and 4000 Hz) and HFA (average in dB HL at 4000, 6000 and 8000 Hz)

Participant	Age	Sex	ΡΤΑ	HFA	Lesion
l (1)	64	М	22.5	28.1	Paramedial right thalamus
I (9)	32	М	15	11.6	Right insula infarct
I (4)	52	Μ	25	26	Left medulla oblongata, occipital lobe, hippocampus and right cerebellum infarct
l (6)	32	М	5.5	6.6	Right temporal lobe infarct
SC (3)	44	М	8.3	11.6	Right putamen/corona radiata infarct
SC (5)	53	F	25	28.3	Right superior parietal lobule infarct
SC (2)	24	М	18.3	16.6	Left frontotemporal and insula infarct
SC (7)	78	М	25	30.1	Left occipito-temporal infarct
SC (8)	64	М	22.5	28.1	Right temporal infarct

A Mann-Whitney U test showed no statistically significant difference for age, PTA and HFA of intervention and the standard care group. dB, decibel; F, female; HFA, high-frequency average; HL, hearing level; I, intervention; M, male; PTA, pure-tone average; SC, standard care.

- ▶ a PTA;
- ► auditory processing tests (speech in babble test,¹⁵ gaps in noise test,¹⁶ perceptual property, apperceptive and semantic processing tests¹⁷);
- the WAB; 18
- ▶ the AIAD questionnaire.¹⁹

Outcome assessment tools, testing methods and outcome measures

Outcome assessments were conducted during visit 1 within 1 week from the baseline assessments, and visit 2 at 10 weeks later. The outcome assessment tool was *the Bamford-Kowal-Bench* (*BKB*) *sentence* $test^{20}$ presented against a 20-talker babble noise, conducted within the 'crescent of sound' booth.²¹ Repetition of at least three keywords per sentence is required for correct performance. The level of the sentences and the background noise are adaptively varied to estimate the speech reception threshold (SRT) which is the SNR for 50% correct performance.

The AB-York Crescent of Sound is a sound-attenuated booth with audio stands in a semicircular arc at a 1.45° m radius from the participant's chair.²¹

The BKB sentences were presented from the loudspeaker positioned at 0° azimuth to the participant with the babble coming:

- A. from the same loudspeaker at 0°at the front;
- B. from the loudspeaker 90° to the left (-90°) ;
- C. from the loudspeaker 90° to the right (+ 90°).

The test was conducted with the participant using the FM (aided condition) or not using the FM (unaided condition). Testing for each of the three locations was repeated twice. Each participant completed 12 runs of the sentences in noise test as follows:

- 1. Aided condition: The FM transmitter microphone was positioned on a stand 12 cm in front from the 0° azimuth loudspeaker and the participant wore the personal binaural FM systems in their ears. Two runs of the test were conducted for the babble noise coming from each of the three locations: straight-ahead (0° azimuth), left (-90°) and right ($+90^{\circ}$) loudspeaker.
- 2. Unaided condition: The same protocol as per the aided condition with the babble noise coming from each of three locations ×2 runs, but with the participant not wearing the binaural FM systems.

The order of the tests was done randomly across participants. A different sentence list was used in each test run.

The primary outcome measure for this test was the SRT obtained for speech and noise presented from 0° (S0°N0°), speech presented at 0° and noise 90° to the left (S0°N–90°) and speech presented at 0° and noise 90° to the right (S0°N+90°).

Intervention

Intervention subjects

At visit 1, all four intervention subjects were provided with the Phonak iSense Micro receiver and the ZoomLink+ transmitter. The FM receiver gain is varied proportionally to the noise level at the microphone of the FM transmitter. The default FM receiver gain is set to +10 for noise up to 57 dB sound pressure level (SPL) and with a maximum gain of +24 for noise of \sim 75 dB SPL.²² When speech is not present at the FM microphone, the receiver is muted.

Intervention subjects were instructed how to check function of the system and asked to change batteries every 2 weeks. They were advised to use the FM system for 7 days a week and about 6 hours daily over the next 10 weeks. Patients were asked to use the FM at home with family members and with multiple media devices such as music players, radio, television and their computer. They were also asked to use the FM in social situations such as family outings (with their significant others wearing the microphone), restaurants, public houses as well as small group meetings at work and so on. Written instruction was provided together with standard listening strategies handout.

Standard care subjects

At visit 1, all standard care subjects were given an oral explanation of their listening difficulties and a standard listening strategies advice handout. They were retested at visit 2, 10 weeks later from visit 1.

Statistical analysis

All analyses were carried out using STATA 11. One-way analysis of variance (ANOVA) and analysis of covariance (ANCOVA) were conducted. In addition, bootstrapping procedures, which facilitate parametric statistical analyses on non-normally distributed data sets, were used to address the limitation of the small sample size. Such procedures estimate statistical parameters based on a large number of random samples (with replacement) from an original data set. Bootstrapped ANOVA and ANCOVA were thus conducted²³²⁴ to determine a statistically significant difference between the intervention and standard care groups on the SRTs in noise at the baseline time point and at 10 weeks. Bootstrapped z-statistic and p values (bias corrected, accelerated with 1000 replications) were calculated. ANCOVA was performed to control for age and side of lesions, and to examine the differences between the intervention and standard care groups on postintervention scores, while controlling for baseline scores of the same measure. The independent variable, study group, included two levels: intervention group and standard care group. The dependent variable was the SRT in noise scores and covariates were age and side of lesion.

A Mann Whitney test was used to compare the primary outcomes between the intervention and standard care groups at baseline and 10 week time point.

RESULTS

There were no drop outs to the study, and all recruited subjects came back for retest 10 weeks later. All four

intervention subjects complied with the daily use of the FMs. According to their own reports the FMs were used for a minimum of 4 hours every day.

The simple and bootstrapped one-way ANOVA showed no statistically significant difference between the intervention and standard care groups on the SRTs in noise at visit 1 time point. There was no difference between groups with respect to the SRTs in noise when the babble was presented at either the left or the right side, in aided and unaided condition (tables 2 and 3). A series of simple and bootstrapped univariate ANOVAs were calculated to examine the differences between the intervention and comparison groups on visit 2 (post intervention) scores. Simple and bootstrapped ANOVA showed a statistically significant improvement in SRT in noise when the noise was coming from left and right for aided and unaided conditions at visit 2 time point in the intervention group compared with those who received standard care. However, when a series of simple and bootstrapped univariate ANCOVA were calculated to control for baseline outcomes, age and side of lesions, a statistically significant improvement in SRT in noise was observed only when the noise was coming from left for aided and unaided conditions at visit 2 time point in the intervention group compared with those who received standard care (tables 2 and 3).

Figure 1 shows the SRT individual scores for intervention and standard care subjects at visit 1 and visit 2, in aided and unaided conditions.

A series of Mann-Whitney U tests were performed in the different speaker-testing positions, these were for baseline visit and 10-week visit to assess differences in the primary outcomes of SRT in noise between intervention and standard care groups. The results are summarised in table 4.

DISCUSSION

Forty-four per cent of eligible stroke subjects for the FM intervention (ie, with preserved peripheral hearing but

with speech in noise reported difficulties and test deficits) were willing to use the FM systems daily. In addition to the immediate benefit in terms of improved speech in noise reception threshold in the aided condition (with FM system), the intervention subjects showed a further substantial benefit for speech in noise test performance even in the unaided condition after a period of 10 weeks daily FM use in the versus no benefit in the standard care subjects. The benefit exceeded 6 dB which is the cut-off value for patients to clinically seek such intervention⁸ when the noise was on their left side. The study has small case numbers, and findings ought to be interpreted with caution. Low study numbers affect precision of measurements, and even a nominally statistically significant finding may not reflect a true effect.²⁵ Furthermore, groups were not randomly allocated, thus being prone to selection bias, while willingness for randomisation and the factors influencing patient preference and thus group allocation²⁶ were not determined. An additional limitation was that we could not investigate the effect of extent of brain lesion on results. However, the study purpose was to determine if there is potential value in running a large-scale intervention trial.

Taking the methodological caveats into consideration, it would be useful to consider the study's findings implications. Improved SRTs when using the FM system after prolonged daily FM use could be attributed to acclimatisation-type effects (ie, getting used to the device effects), such as improved perception of highfrequency phonemes over time.²⁷ However, such benefits—if any—tend to be small.²⁸ The findings of a robust improvement in SRTs in the unaided condition in the FMs intervention group may thus indicate that a brain plasticity mechanism is involved.

In the presence of normal pure-tone audiometric thresholds and in the absence of aphasia, our stroke subjects had degraded speech encoding (as indicated by their abnormal speech in babble test results) and selfreported difficulties with speech in noise due to their-

Table 2 ANCOVA and ANOVA. Intervention versus standard care									
	ANCOVA and ANOVA Intervention vs standard care								
		MS		F-statistic		p-Value			
Condition		ANCOVA	ANOVA	ANCOVA	ANOVA	ANCOVA	ANOVA		
S0N-90 unaided	Visit 1	9.6	0.13	1.1	0.01	0.45	0.91		
	Visit 2	47.8	140.4	5.5	21.2	0.04*	0.002*		
S0N+90 unaided	Visit 1	5.1	14.0	1.4	5.2	0.33	0.06		
	Visit 2	12.9	37.2	3.1	11.6	0.13	0.01*		
S0N-90 aided	Visit 1	5.9	0.12	0.6	0.01	0.60	0.90		
	Visit 2	57.76	149.2	5.2	13.1	0.05*	0.008*		
S0N+90 aided	Visit 1	3.6	8.2	0.3	0.89	0.83	0.37		
	Visit 2	42.5	121.1	1.88	7.1	0.24	0.03*		

The analysis of covariance (ANCOVA) model included age and side of stroke as covariates.

ANOVA, analysis of variance; MS, mean square.

*(in bold) statistically significant at p at or < 0.05

Table 3 Bootstrapped ANCOVA and ANOVA. Intervention versus standard care **Bootstrapped ANCOVA and ANOVA** Intervention vs standard care Bootstrapped Bootstrapped Bootstrapped z-statistic SE p-value ANCOVA Condition ANCOVA ANOVA ANOVA **ANCOVA** ANOVA S0N-90 unaided Visit 1 4.4 2.5 0.1 -0.02 0.9 0.9 Visit 2 -4.4 0.05* 0.000* 4.1 1.7 -1.8 S0N+90 unaided Visit 1 -2.2 4.1 1.1 -0.6 0.6 0.07 0.002* Visit 2 8.1 2.4 -0.5 -3.2 0.6 S0N-90 aided Visit 1 6.6 1.9 -0.2 0.13 0.8 0.9 Visit 2 4.9 1.1 -2.0 -6.8 0.04* 0.000* S0N+90 aided Visit 1 5.5 1.8 -0.2 -1.05 0.8 0.3 Visit 2 2.3 0.02* 8.7 3.3 -0.9 0.3

The analysis of covariance (ANCOVA) model included age and side of stroke as covariates.

ANOVA, analysis of variance.

*(in bold) statistically significant at p at or < 0.05

Figure 1 (A and B) Speech reception threshold (SRT) at visit 1 and visit 2 in aided (A) and unaided conditions (B) for noise coming 90° to the left and for noise coming 90° to the right. SRT, speech reception threshold; S, subject.

4. Left aided condition visit 1 Left aided condition visit 2 0 SRT in noise -4 -8 12 Right aided condition visit 1 -16 Right aided condition visit 2 -20 -24**±** Intervention S1-Intervention S2-Intervention S3-Intervention S4-Standardcare S2 Standardcare S3 Standardcare S4 Standardcare S5 Standardcare S1

B Individual SRT in noise scores [unaided condition]



A Individual SRT in noise scores [aided condition]

		Intervention		
Condition	Group	Median (variance)	Standard care	p-Value
S0N–90 unaided	Baseline	1.1 (13.9)	-1.4 (2.8)	0.81
	10 week	-7.4 (5.8)	-0.15 (7.2)	0.01*
S0N+90 unaided	Baseline	-1.4 (2.8)	0.9 (2.6)	0.054
	10 week	-2.5 (4.8)	1.3 (2.5)	0.01*
S0N–90 aided	Baseline	-10.5 (2.8)	-10.6 (13.9)	1.0
	10 week	-20 (7.8)	-11.6 (14.0)	0.01*
S0N+90 aided	Baseline	-10.3 (12.1)	-8.7 (7.1)	0.32
	10 week	-16.2 (29.6)	-8.9 (7.5)	0.05

stroke related auditory processing deficits as evidenced by their abnormal results in other non-speech auditory processing tests. This resulted in impaired identification of lexical items in stored knowledge. Additionally, the increased effort required for speech discrimination because of background noise would be expected to reduce their information processing capacity and thus the short-term memory required for speech recall and understanding.²⁹ It has been proposed³⁰ that each level of the auditory cortical hierarchy attempts to predict the sensory representation of the speech signal of interest at the level below by transmitting a top-down prediction, with prediction error information (ie, the incoming signal differing from the prediction) transmitted back to the higher level, leading to a recalibration of the higher level representations. This updating of predictions at higher level is heavily dependent on attention mechanisms influenced by the degree of salience of the stimulus and the listening context.³¹ It is thus postulated that in our patients with stroke, auditory processing deficits reduced the clarity of incoming speech and the brain (attention) responds by decreasing the sensory precision or postsynaptic gain. This was to some extent redressed when stroke subjects used the FM system within the crescent of sound, as demonstrated by the immediate improvement in SRTs in the aided versus the unaided condition in visit 1. It may further be postulated that prolonged use of the FMs over 10 weeks, with active listening to salient speech (assuming that cases used the FM system to listen to speech of interest to them) led to attention optimising the synaptic gain that represents the precision of the bottom-up sensory information (prediction error) during the hierarchical inference process within the auditory brain. Candidate brain regions where this effect could take place include primary auditory cortex area, or Broca's and inferior frontal gyrus, that is, the areas showing a relative specialisation for phonological encoding³² or subcortical areas, in accordance with the results of Hornickel *et al.*¹⁰

This is the first study to investigate potential auditory plasticity following a prolonged use of FM systems in adult patients with acquired stroke brain lesions. However, this study has methodological limitations that do not allow to draw definitive conclusions. The findings presented here should motivate further work with a larger randomised clinical trial to investigate the true effect size of prolonged FM-use benefits, the effect of type and extent of brain lesion and of cognitive and linguistic factors, as well as the mechanism for any benefits. Further research is required to look into this promising intervention that may benefit $\sim 17\%$ of the stroke population.⁷

Author affiliations

¹UCL Ear Institute, London, UK

²National Hospital for Neurology and Neurosurgery, London, UK

³UCL Speech Hearing and Phonetic Sciences, London, UK

⁴UCL Dementia Research Centre

⁵Department of Brain Repair and Rehabilitation, Stroke Research Centre, UCL Institute of Neurology

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