

Benefits of Cochlear Implantation in Middle-Aged and Older Adults

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Christiane Völter 
Lisa Götze
Imme Haubitz
Stefan Dazert
Jan Peter Thomas

Department of Otorhinolaryngology,
Head and Neck Surgery, Katholisches
Klinikum, Ruhr-University of Bochum,
Bochum, Germany

Introduction: Nowadays cochlear implantation (CI) is the treatment of choice in adults in case conventional hearing devices fail. Besides speech perception, an improvement in quality of life and in cognitive performance has been reported. Thereby, the study focused on the impact of age. **Participants and Methods:** Thirty middle-aged (MA) between 50 and 64 years and 41 older subjects (OA) aged 65 and older with bilateral severe hearing loss performed a comprehensive computer-based neurocognitive test battery (ALAcog) pre- and 12 months post-implantation. Besides, monosyllabic speech perception in quiet (Freiburg monosyllabic speech test), health-related quality of life (HR-QoL, Nijmegen Cochlear Implant Questionnaire) and depressive symptoms (GDS-15) have been assessed.

Results: Both age groups significantly improved in all three categories after 12 months. No differences were evaluated between MA and OA regarding speech perception and HR-QoL pre- and post-operatively. In contrast, cognitive performance differed between the age groups: pre-operatively OA performed worse in most neurocognitive subdomains like working memory ($p=0.04$), inhibition ($p=0.004$), processing speed ($p=0.003$) and mental flexibility ($p=0.01$), post-operatively MA outperformed OA only in inhibition ($p=0.01$). Age only slightly influenced cognitive performance in MA, whereas in OA age per se tremendously impacted on working memory ($p=0.04$), inhibition ($p=0.02$), memory ($p=0.04$) and mental flexibility ($p=0.01$). Educational level also affected processing speed, mental flexibility ($p=0.01$) and working memory ($p=0.01$). This was more pronounced in OA. In both age groups, hearing status had a strong effect on attentional tasks ($p=0.01$). In MA, depressive symptoms were more influential on cognitive functioning and on HR-QoL than in OA. Improvement in quality of life ($p=0.0002$) and working memory ($p=0.001$) was greater for those with a higher pre-operative depression score.

Conclusion: Speech perception and HR-QoL improved in hearing impaired, independently of age. Pre-operative differences in cognitive performance between OA and MA clearly attenuated 12 months after CI. Impact of comorbidities differed between age groups.

Keywords: cochlear implantation, age-related hearing loss, benefit, outcome, cognitive domains, quality of life, depression

Introduction

The growing number of the elderly population will be one of the major challenges in the future. Health-related disorders will become an increasing problem for the individual and a large socio-economic burden for the general population. Multiple chronic diseases including deficits in sensory abilities show a highly rising number. According to the World Health Organization (WHO) already now more than 450 million people worldwide suffer from disabling hearing loss and more than 50% of octogenarians are hereby affected.¹

Correspondence: Christiane Völter
Tel +49 2345098390
Email christiane.voelter@rub.de

Recently consequences of age-related hearing loss have gained further interest.²⁻⁵ Due to changes in communication patterns, quality of life is strongly reduced in hearing-impaired subjects.^{2,6} According to the National Council on the Aging Report comprising 4394 participants only 39% of hearing-impaired elderly have an excellent health compared to 68% in the general population.⁷

The reduced quality of life due to hearing loss has been largely described.^{2,8,9} In the Blue Mountain Hearing Study conducted by Chia who enrolled 2431 participants the physical and mental domain of the self-reported quality of life assessed by the Short Form-36 Questionnaire was associated with the severity of the hearing impairment and the use of hearing aids positively influenced outcome.¹⁰

Hearing-impaired subjects have a higher risk to suffer from social isolation or to fall into depression.^{11,12} This applies especially to women: female sexagenarian with a hearing loss of at least 25 dB have a 3.5-fold increased risk to suffer from isolation in comparison to age-matched normal hearing controls.⁵ This is in line with large epidemiologic longitudinal studies by Li et al analyzing data of more than 18,000 adults in the US and by Hsu et al on more than 20,000 subjects in Taiwan^{11,13} and with a recently published review and meta-analysis including a total of 147,148 participants from 35 studies. Hereby, hearing loss was associated with a significantly greater odds ratio for depression in older adults (OR = 1.479).¹⁴ Interestingly, mainly minor depressive symptoms and not severe depression seem to be highly associated with hearing impairment in the long-term follow-up as reported by Cosh in a study on 8340 adults.³ This might be even more pronounced in adults aged <65 years.^{15,16} Besides the negative impact of age on sensory abilities, aging is associated also with an enhanced cognitive decline starting already from the second decade of life.¹⁷ Further, executive functions decrease from 4% in the fourth decade to 11% for adults in their seventh decade as shown by Singh-Manoux et al in 4675 elderly.¹⁸ Retirement negatively impacts on cognitive functioning (such as memory) as described by Bonsang 2012 in a large data set of 82,462 elderly aged 51–75 years.¹⁹ Whereas fluid intelligence such as inhibition, interference control and working memory is highly vulnerable to age, crystalline intelligence persists mostly stable or even increases in ongoing life.^{20,21}

The multisensory impact on cognition has been extensively studied by Humes in 2013 in 98 hearing-impaired persons with a mean age of 69.2 and a control group of normal hearing participants aged 22.7 years. Younger subjects outperformed the elderly in working memory and

verbal processing speed. In contrast, elderly achieved better results in tests with a high context portion.⁶ In addition, sensory-cognitive associations were stronger if more than one sensory domain was affected.²²

Thereby hearing and cognition cannot be considered as variables separately affected by age, but they are closely related and influenced by each other. Cognitive functioning is required to ensure speech perception especially in adverse listening conditions on one side.^{6,23,24} On the other hand, poorer hearing is associated with a worse cognitive performance, and hearing is proposed as one modifiable risk factor, which might diminish the risk of dementia if treated in the mid-life.^{25,26}

However, psychological disorders as depression also have an effect on cognition and might be important mediators in the relationship between hearing loss and cognitive decline in the elderly.^{27,28} This has already been described by Huber in a study on 30 normal hearing (NH) and 30 hearing-impaired (HI) subjects between 60 and 80 years.²⁹ HI subjects significantly differed in their performance in the clock drawing test, the word list learning (immediate and delayed) as well as in the Stroop and the TMT B tasks. In terms of anxiety assessed by the HADS (Hospital Anxiety and Depression Scale) no difference was obtained between the NH and HI subjects, but HI presented significantly higher levels of depressive symptoms. Furthermore, TMT B performance was mediated by the severity of depressive problems, whereas none of the other cognitive subtests did.

Nowadays, cochlear implantation has become the treatment of choice in severe to profound hearing-impaired subjects, which cannot be sufficiently treated by conventional hearing devices.³⁰ Multiple studies have shown that also the older population has a substantial audiological benefit hereby.³¹⁻³³ However, cochlear implantation is still less frequently performed in the aged hearing-impaired population than it should be done due to the audiological necessity even in industrialized nations as stated by Turunen-Taheri who studied the treatment of 1076 older subjects with severe to profound hearing loss at a mean age of 70.6.³⁴

In general, hearing benefit regarding speech perception in quiet is quite similar between different age groups even if some studies point out that older adults need more time to adapt to the new auditory signal or do not reach the same speech perception in noise.³³ Age differences in demanding listening situations might be due to the longer duration of

hearing loss or to the influence of cognitive abilities, especially inhibitory control and attention. Besides pure audiological benefits also significant improvements in post-operative quality of life have been described. Patients aged 70 years and older even outperform younger adults aged between 19 and 67 in health-related questionnaires such as the Nijmegen Cochlear Implant Questionnaire (NCIQ).³¹

Recently there is an increasing interest in neurocognitive changes in older subjects after hearing rehabilitation.^{35–39} Mosnier was the first who investigated cognitive functions in older hearing-impaired subjects between 65 and 85 years before and after cochlear implantation and reported on benefits mainly regarding recall and attention.⁴⁰ An improvement in attention has also been found by Sarant in 59 CI patients aged 72 that underwent testing with the Cogstate battery 18 months after cochlear implantation.³⁸ Besides, working memory improves post-implantation as shown in a recent study by our group in 20 elderly 12 months after implantation.³⁶ Similar findings have also been described by Jayakody in 16 CI recipients with a mean age of 61.7 that underwent testing with the Cambridge Neuropsychological Test Automated Battery (CANTAB).³⁵

However, the impact of age has not been extensively evaluated in these studies although hearing, age and cognition are closely related to each other. Furthermore, the auditory-presented material used in some cognitive assessments might have partially influenced previous results in severely hearing-impaired subjects.^{41,42} Therefore, the presented study aimed to analyze the impact of age on the benefits of CI 12 months after implantation in a large sample size regarding speech perception and quality of life as well as cognitive performance using a comprehensive non-auditory test battery.

Participants and Methods

Participants

Age groups were defined in middle- (MA) and older-aged (OA) adults according to the definition of the WHO. Middle-aged was classified as people aged between 50 and 64 years, and old age starts at 65 years (WHO 2002) when individuals usually retire. Thirty MA and 41 OA with a bilateral severe to profound postlingual hearing loss were included (Table 1). The two groups did not differ with regard to gender ($p=0.78$), education ($p=0.78$) nor with regard to the duration of severe hearing impairment ($p=0.74$) or the amount of depressive symptoms ($p=0.43$). Inclusion criteria were defined as (1) age of 50 or older, (2) native or excellent

German language speaker, (3) no uncorrected vision loss, (4) absence of a global cognitive impairment according to the Multiple Word Sentence Test (MWT-B),⁴³ (5) no history of severe depression, (6) absence of central nervous system disease or treatment with anticholinergic medication, (7) postlingual bilateral hearing loss of 61dB or worse on average in the frequencies 0.5, 1, 2 and 4kHz on the better hearing ear.

All participants were recruited at the ENT-department at the Ruhr-University Bochum, Germany.

Audiometric Assessment

Pre-operatively, pure tone audiometry was measured separately for each ear at the frequencies between 0.25 and 8 kHz (DIN EN ISO 8253). The 4-pure tone average (4-PTA) was calculated separately for each ear using the mean of the frequencies 0.5, 1, 2 and 4 kHz.

The German Freiburg monosyllabic speech test in quiet was performed in all subjects pre- and 12 months post-operatively in free field at 65 dB sound pressure level (SPL) by an experienced audiologist in a soundproof booth. Additionally, in 37 patients, sentence recognition in noise was assessed 12 months postoperatively by the Oldenburg sentence test.⁴⁴ For speech perception in noise, the signal-to-noise ratio (SNR) for 50% correct word recognition was assessed. An adaptive procedure was

Table 1 Demographic Data

	Middle-Aged Recipients	Older-Aged Recipients
Number of participants	n= 30	n= 41
Mean age in years (SD)	Total group 57.33 (4.48); range 50–64	Total group 72.33 (5.27); range 65–84
	Male n= 12 57.3 (4.68); range 50–64	Male n= 12 72.92 (5.82); range 65–84
	Female n= 18 57.3 (3.56); range 50–64	Female n= 29 73.0 (5.08); range 65–81
Mean educational background in years (SD)	Total 12.03 (2.15) Male 12.00 (2.00) Female 12.05 (2.29)	Total 12.15 (2.42) Male 12.41 (2.97) Female 12.03 (2.20)
Duration of severe hearing impairment in years (SD)	23.59 (15.95); range 1–55	22.26 (13.73); range 1–57

Note: SD indicates standard deviation.

used with background noise fixed at 65 dB SPL and both speech and noise were presented from the front. A lower score indicated a better sentence comprehension in noise.

Health-Related Quality of Life

The Nijmegen Cochlear Implant Questionnaire (NCIQ) was used pre- and post-operatively to assess health-related quality of life (HR-QoL).⁴⁵ Patients evaluated quality of life in 60 statements covering three domains which were further split into six subdomains: 1) physical domain: (a) basic sound perception and (b) advanced sound perception, (c) speech production; 2) psychological domain: (a) self-esteem; 3) social domain: (a) activity limitations, (b) social interactions. Answers range from 0 (not at all) to 100 (very good). A higher score reflected better health-related quality of life.

Neurocognitive Assessment

A visually based neurocognitive assessment tool (ALAcog), which is based on standardized paper and pencil versions adapted for hearing-impaired and already used in former studies, has been applied pre- and post-operatively.^{46,47} The test battery is composed of 11 subtests, covering various cognitive subdomains as previously described in detail.^{36,47}

-M3 (according to the D2 test of attention by Brickenkamp 1962)⁴⁸ to assess attention.

-Recall and Delayed Recall (according to the verbal learning and memory test by Helmstaedter 2001)⁴⁹ for short and delayed memory.

-0- and 2-back (based on Kirchner 1958)⁵⁰ to assess working memory.

-OSPAN (based on the Operation Span task by Conway 2005)⁵¹ is a dual task, which assesses working memory as well.

-Flanker (according to Eriksen and Eriksen 1974, Wild-Wall 2008)^{52,53} measures the ability to inhibit compatible (cFlanker) or incompatible (iFlanker) distractors.

-TMT A and B (based on the Trail Making Test by Reitan 1958).⁵⁴ TMT A assesses processing speed, TMT B mental flexibility.

-Verbal fluency (according to the Chicago Word Fluency Test by Thurstone 1948)⁵⁵ assesses verbal functioning and executive control.

For each test raw data and a total score, the inverse efficiency (IE) was calculated based on the time needed and the number of correct answers given.⁵⁶ A better cognitive performance is indicated by a lower IE score. The

bias of practice effects was minimized by the application of two different versions.

Psychosocial Comorbidities

Severe depression has been ruled out in all patients by precise questioning. Besides in 53 out of 71 subjects with a mean age of 65.87 (SD 9.34), the GDS-15-Assessment (Geriatric Depression Scale) was applied pre-operatively. This test is based on 15 dichotic questions on the attitude and mood of the subjects. A score of 0–5 indicates the absence of depressive symptoms, 6–10 indicates mild or moderate depressive symptoms and a score >10 might be a hint for profound depressive symptoms.⁵⁷

Educational background has been assessed by the number of years participants attended school or did further studies.

Aural Rehabilitation

All participants took part in a standardized regular aural rehabilitation program once a week during the first weeks post-implantation (basic rehabilitation) with an experienced speech therapist in a face-to-face session in accordance with the guidelines of the German Society for Otorhinolaryngology, Head and Neck Surgery (Guidelines for cochlea implant supply, AWMF). After this, aural training was followed up to 2 years at a lower frequency.

Statistical Analysis

Data analysis was performed in Medas (C. Grund, Margetshoechheim, Germany). All results were presented using mean and standard deviation (SD). As dataset was partially not normally distributed, Wilcoxon- and Mann-Whitney *U*-test were applied to compare cognitive and auditory abilities as well as quality of life pre- and post-operatively. TMT was calculated with linear models and rule of proportion in case participants were unable to finish the task within 90 seconds.

The effect size (EF) has been calculated adapted to Cohen's *d* (up to 0.1 refers to a weak, 0.3 to a moderate and 0.5 to a strong effect). To address, whether speech perception at 65dB, age, gender or education were most predictive for cognitive performance 12 months post-implantation, multiple regression analysis was performed. Standardized Beta-weight (β) was used to compare the predictors.

Correlation analyses using Kendall's tau (τ) were performed between different variables such as speech perception scores, quality of life measures, depressive symptoms,

cognitive function and duration of deafness. The level of significance was set to $p < 0.05$.

The study was in line with the requirements of the ethic institution of the Ruhr University Bochum (No 16–5727-BR) and the requirement of the declaration of Helsinki. All participants signed their informed consent.

Results

Hearing Status

Preimplantation both age groups did not significantly differ with regard to 4-PTA of the worse or the better hearing ear ($p=0.07$, $p=0.9$). Moreover, speech understanding before implantation was comparable between the groups without any significant difference ($p=0.68$). Twelve months after implantation patients of both age groups showed a significant improvement of monosyllabic speech perception (MA $p=0.003$ and OA $p=0.005$) with similar results regarding the speech perception scores in MA and OA recipients ($p=0.45$) (Table 2). Also, 12 months after implantation speech perception in noise was similar ($p=0.55$) in MA and OA.

Correlation Analysis

No correlation was evaluated between speech perception in quiet and total quality of life. The only significant association found was with regard to the post-operative subscore self-esteem, but only in the elderly. Subjects who improved more in speech perception at 65 dB showed a significant better self-esteem ($\tau=0.25$; $p=0.03$). However, speech perception in quiet as well as improvement in speech perception in quiet after 12 months was predicted by pre-operative inhibition of compatible stimuli ($\tau=-0.22$, $p=0.049$; $\tau=-0.24$; $p=0.04$, respectively).

Health-Related Quality of Life

Age had a partial influence on pre-operative quality of life. Older subjects scored better in basic sound perception ($p=0.047$). Twelve months post-implantation both groups revealed a significant improvement of the HR-QoL ($p=0.0000$) with similar total scores of the NCIQ between both age groups pre- and post-implantation (pre-implantation MA 46.02 vs OA 50.55; $p=0.5$ and post-implantation MA 65.33 vs OA 66.42; $p=0.69$) (Table 3).

Correlation Analysis

Twelve months post-operatively, no significant correlation could be detected between speech perception in quiet and total NCIQ-Score ($p=0.19$), but with regard to the

Table 2 Results of Hearing Assessment

CI-Recipients	Middle-Aged (MA)	Older-Aged (OA)	p
Pre-implantation			
4-PTA worse ear (mean)	101.12dB	96.37dB	0.07
4-PTA better ear (mean)	79.33dB	79.23dB	0.99
Freiburg monosyllabic speech perception at 65 dB SPL (mean)	5.3%	7.5%	0.68
Post-implantation			
Freiburg monosyllabic speech perception at 65 dB SPL (mean)	60.88%	56.33%	0.45
Speech perception in noise (mean)	+0.8dB	+1.26dB	0.55

Notes: SPL indicates sound pressure level. Significance was set to $p < 0.05$.

improvement of quality of life in MA and OA (MA $\tau=-0.28$; $p=0.048$ and OA $\tau=0.23$, $p=0.04$). However, post-operative speech perception in noise correlated directly to the subdomains of advanced sound perception ($\tau=-0.42$; $p=0.04$) and of self-esteem ($\tau=-0.31$; $p=0.02$) in MA, but not in OA subjects.

Duration of deafness correlated only pre-operatively with NCIQ score in some subdomains such as in self-esteem ($\tau=0.30$, $p=0.03$) in MA as well as in the physical domain ($\tau=0.23$, $p=0.04$) and advanced sound perception in OA ($\tau=0.22$, $p=0.04$). Post-operatively, there was no association between duration of hearing loss and quality of life ($\tau=-0.002$, $p=0.98$). However, older subjects showed a larger improvement in post-operative advanced sound perception in case of a shorter duration of hearing loss ($\tau=-0.22$, $p=0.04$).

Neurocognitive Performance

In total, a significant improvement could be observed in cognitive functioning post-operatively in the whole study group with the largest improvement in patients with poor baseline cognitive performance.

Pre-operatively OA subjects performed worse than MA in 4 out of 11 neurocognitive subtests. This was highly significant with regard to the TMT (TMT A $p=0.003$, $EF=0.61$ and TMT B $p=0.01$, $EF=0.56$ and the inhibition (iFlanker $p=0.004$, $EF=0.39$). Furthermore, significant age-dependent differences could be detected in working

Table 3 Health-Related Quality of Life Assessed by the Nijmegen Cochlear Implant Questionnaire (n=71)

Subtest		Pre-Implantation			Post-Implantation		
		Mean	SD	p	Mean	SD	p
Total	MA	46.02	15.32	0.5	65.33	13.63	0.69
	OA	50.55	16.43		66.42	15.26	
Physical	MA	48.88	18.53	0.13	68.87	13.32	0.31
	OA	56.58	17.26		71.93	15.44	
Social	MA	43.05	17.97	0.44	61.42	18.67	0.93
	OA	42.19	19.56		60.97	19.73	
Basic sound perception	MA	41.21	20.55	0.047 *	68.93	15.95	0.41
	OA	52.98	21.92		71.39	17.81	
Advanced sound perception	MA	39.99	22.59	0.19	62.54	18.77	0.66
	OA	47.7	22.58		64.44	20.18	
Speech production	MA	65.44	21.64	0.69	75.13	16.64	0.18
	OA	67.73	16.41		79.94	14.00	
Self-esteem	MA	43.36	16.25	0.5	62.53	16.95	0.41
	OA	47.19	17.62		60.77	14.48	
Activity limitations	MA	42.13	18.78	0.91	60.49	19.29	0.9
	OA	42.99	20.59		60.03	20.68	
Social interactions	MA	43.98	18.92	0.29	62.34	19.64	0.95
	OA	41.3	19.93		61.91	20.75	

Notes: MA indicates middle-aged participants aged between 50 and 64 years and OA includes an age of ≥ 65 years. *Means significant ($p < 0.05$).

memory as assessed by the OSPAN ($p=0.04$, $EF=0.53$). The effect size was between 0.39 and 0.61. Post-operatively, MA outperformed OA still in the ability to suppress inhibitory signals (iFlanker $p=0.01$), with a small effect size of 0.23 (Table 4). MA and OA improved similar in most subtests studied. Only in 0-back ($p=0.04$) as well as in TMT A ($p=0.02$) improvement was more pronounced in OA than in MA.

In-Depth Analysis

Pre-operatively a slower reaction time and less correct items were found for OA than for MA subjects in the M3 task, but without any significance (reaction time MA 840.64 ms vs OA 945.65 ms; $p=0.11$ and correct letters MA 79.9 vs OA 73.6; $p=0.16$). Post-implantation the two groups significantly differed: OA completed only 85.78 trials, whereas MA were able to increase the performance to 95.77 trials ($p=0.02$). In addition, the number of correct labels was higher for MA than for OA ($p=0.046$) as well as the reaction time was shorter for the younger than for the older group (MA 759.41 ms vs OA 852.46 ms; $p=0.03$).

With regard to Recall and Delayed Recall, pre- and post-operative results were independent of the age groups. Patients of both age groups scored less in Delayed Recall than in Recall tasks by more than 1.5 words on average. Both groups equally improved pre- to post-implantation (Recall pre: $p=0.91$ vs post: $p=0.12$ and Delayed Recall pre: $p=0.37$ vs post: $p=0.08$).

The same was true for the 0- and the 2-back task; 0-back performance was comparable between OA and MA pre- and post-implantation (pre: MA 342.30 vs OA 324.53; $p=0.79$ and post: MA 314.16 vs OA 337.07; $p=0.12$).

Working memory assessed by the OSPAN significantly differed between OA and MA ($p=0.04$, $EF=0.53$) pre-operatively. This was due to the smaller number of correctly solved maths' calculations in the OA (MA 38.96 vs OA 37.68; $p=0.04$). Post-operatively, OA required longer to solve the equations (MA 2734.27 ms vs OA 3244.22 ms; $p=0.04$), but they calculated as well as the MA (correct equations: $p=0.93$).

A highly significant difference could be detected between the two age groups in the pre-operative TMT

Table 4 Pre- and Post-operative Neurocognitive Performance According to Age (n=71)

Subtest	Age Group	Pre-Implantation				Post-Implantation			
		Mean	SD	p	Effect Size	Mean	SD	p	Effect Size
M3	MA	930.50	343.32	0.11	0.31	806.1	370.49	0.06	0.57
	OA	1083.78	577.22			908.78	324.54		
Recall	MA	529.00	190.19	0.91	0.01	429.66	197.78	0.12	0.08
	OA	526.09	203.05			489.02	204.98		
Delayed Recall	MA	667.66	166.85	0.37	0.13	558.00	232.23	0.08	0.20
	OA	692.92	207.07			646.82	225.18		
0-back	MA	342.30	106.56	0.79	0.22	314.16	37.10	0.12	0.18
	OA	324.53	57.37			337.07	61.56		
2-back	MA	899.16	1251.88	1.0	0.27	613.33	255.71	0.48	0.13
	OA	675.17	272.30			726.62	914.79		
cFlanker	MA	363.93	64.03	0.08	0.39	363.60	105.98	0.07	0.41
	OA	460.53	323.85			399.10	104.02		
iFlanker	MA	503.73	185.09	0.004 *	0.39	545.90	493.52	0.01 *	0.23
	OA	940.75	1456.07			527.20	143.71		
OSPAN	MA	545.43	239.14	0.04 *	0.53	487.13	223.40	0.08	0.39
	OA	694.63	312.45			584.51	240.42		
TMT A	MA	686.44	291.98	0.003 *	0.61	781.89	437.82	0.16	0.65
	OA	1183.51	1037.81			936.73	631.52		
TMT B	MA	1126.55	621.17	0.01 *	0.56	1284.48	776.31	0.15	0.59
	OA	1541.92	810.14			1521.07	967.16		
Verbal fluency	MA	810.33	61.48	0.75	0.13	750.83	122.35	0.89	0.02
	OA	799.87	88.54			750.73	109.93		

Notes: MA indicates middle-aged participants aged between 50 and 64 years and OA includes an age of ≥ 65 years. *Means significant ($p < 0.05$).

(TMT A, $p=0.003$; TMT B, $p=0.015$) but no longer after implantation (TMT A, $p=0.16$; TMT B, $p=0.1$). Pre-operatively, the last label which has been clicked for TMT A was number 23 by MA and 20 by OA ($p=0.02$). Twelve months after CI MA achieved an equal result whereas OA improved by three additional letters ($p=0.5$). The pre-operative performance of TMT B significantly differed (MA 20 vs OA 16), after implantation the numbers were similar between the two age groups ($p=0.15$).

The ability to detect incompatible stimuli (Flanker task) was highly influenced by age pre- as well as post-operatively. OA missed a higher number of incompatible arrows pre- and post-implantation (pre: MA 1.33 vs OA 3.53; $p=0.04$ and post: MA 1.56 vs OA 1.52; $p=0.04$). OA responded slower to the target than MA although not significantly (pre: MA 410.34 vs OA 448.92; $p=0.11$ and post: MA 408.38 vs OA 442.93; $p=0.01$). In contrast, cFlanker responses were similar

for both groups (pre: MA 363.93 vs OA 460.53; $p=0.08$ and post-implantation MA 363.60 vs OA 399.10; $p=0.07$). The number of missed compatible Flankers equally decreased pre- to post-implantation for both groups (pre: MA 0.80 vs OA 1.53; $p=0.33$ and post: MA 0.33 vs OA 0.95; $p=0.21$). Reaction time remained stable pre- to post-implantation in both age groups (pre: MA 342.16 versus OA 344.71; $p=0.15$ and post: MA 350.61 vs OA 369.65; $p=0.11$).

Verbal fluency did not show any difference related to age-groups (pre: $p=0.75$ and post-implantation $p=0.89$). The number of animals for MA pre-implantation was 6.5 and 6.68 for OA. Post-implantation MA named 7.03 and OA 7.86 correct animals.

Correlation Analysis

Pre-operative inhibitory performance assessed by incompatible Flanker was predictive of post-operative speech

perception in quiet in older subjects, but not in the younger group. (OA: $\tau=-0.22$, $p=0.049$, MA: $\tau=-0.1$, $p=0.48$). In addition, post-operative speech perception in quiet was related to post-operative performance in attentional tasks in OA ($\tau=-0.26$, $p=0.02$ vs MA $\tau=-0.19$, $p=0.16$), whereas improvement of speech perception significantly correlated to an improvement of M3 in both age groups (MA: $\tau=-0.27$, $p=0.048$ and OA: $\tau=-0.23$, $p=0.03$).

Covariants on Neurocognitive Performance

Multiple regression analysis regarding post-operative cognitive performance was calculated separately for MA and OA subjects regarding age, gender, speech perception in quiet and education:

In MA age significantly predicted cognitive performance in 2 of 11 subtests: TMT A ($\beta=0.45$, $p=0.01$) measuring processing speed and M3 evaluating attention ($\beta=0.32$, $p=0.049$). Besides speech perception in quiet had the strongest influence on attentional performance assessed by the M3 ($\beta=-0.47$, $p=0.01$). Longer education times correlated with the results in the Recall ($\beta=-0.39$, $p=0.03$) and OSPAN tasks ($\beta=-0.43$, $p=0.02$), and female gender correlated with better results in verbal fluency ($\beta=-0.4$, $p=0.049$).

In OA age influenced 5 out of 11 subtests: 2-back ($\beta=0.31$, $p=0.04$), Recall ($\beta=-0.3$, $p=0.04$), TMT A ($\beta=0.35$, $p=0.02$), TMT B ($\beta=0.39$, $p=0.01$) and cFlanker ($\beta=0.37$, $p=0.02$). Monosyllabic speech perception also had an impact on attentional tasks (M3 $\beta=-0.4$, $p=0.01$) and on inhibition (iFlanker $\beta=-0.32$, $p=0.04$). Education was predictive for the Recall task ($\beta=-0.3$, $p=0.02$), the OSPAN ($\beta=-0.4$, $p=0.01$), the TMT A ($\beta=-0.38$, $p=0.01$) and the TMT B ($\beta=-0.4$, $p=0.01$).

Comorbidities

Subjects with severe depression were excluded from the study. However, a deeper analysis of 53 subjects revealed that 52.8% suffered from slight or moderate depressive symptoms pre-operatively (mean= 7.32, SD 1.4). Total pre-operative GDS-15 score did not differ according to age (MA 6.25 (SD 2.29) and OA 5.73 (SD 1.36); $p=0.43$) or gender (men 5.77 (SD 2.1), women 6.05 (SD 1.71); $p=0.66$). Pre- and post-operative speech perception as well as improvement of speech perception showed no association to GDS-15 score in both age groups. Duration of hearing loss significantly related to mental health ($\tau=-0.24$; $p=0.01$) in younger, but not in older subjects ($\tau=-0.10$; $p=0.43$).

Depression score had an impact on cognitive performance (1) and quality of life (2). This was more pronounced for younger than for older subjects (Table 5):

(1) Pre-operative depressive score correlated with pre-operative cognitive performance only in subjects aged 50–64 years. This was true with regard to attention ($p=0.03$), working memory (2-back: $p=0.001$ and OSPAN: $p=0.01$), inhibition (cFlanker: $p=0.02$ and iFlanker: $p=0.03$) and TMT (TMT A: $p=0.01$ and TMT B: $p=0.03$).

Post-operative cognitive outcome could be predicted by pre-operative mood mostly in the younger group. Patients reporting on a higher depressive score before implantation also obtained worse cognitive results 12 months post-implantation in incompatible Flanker ($p=0.02$), OSPAN ($p=0.03$) in MA and in TMT A for both age groups (MA $p=0.03$ and OA $p=0.02$).

Besides, subjects aged <65 showed a significantly greater improvement in M3 ($p=0.03$), 0-back ($p=0.01$) and 2-back ($p=0.001$) in case of higher depressive scores. This could not be observed in subjects of ≥ 65 .

(2) Pre-operative mental health correlated to the pre-operative quality of life in both groups, but more pronounced in MA. Post-operative depression score was predictive for health-related quality of life in OA for physical sound perception ($\tau=-0.26$, $p=0.047$). Besides, improvement in quality of life relied on psychosocial comorbidity only in the younger subjects ($p=0.04$).

Discussion

The target of the study was twofold (1) to evaluate the impact of age on cognitive and mental benefits of cochlear implantation in adult cochlear implant recipients and (2) to analyze the age-dependent interaction between speech comprehension, cognition and quality of life.

According to previous investigations also in our study adults of any age achieve a significant improvement in monosyllabic speech perception after cochlear implantation with no differences between the two age groups of middle-aged and older-aged adults.^{33,58–60}

Besides restoration of speech perception, cochlear implantation has also a tremendous impact on the quality of life. In this study quality of life was equal between the age groups at any time despite in the subtest basic sound perception before cochlear implantation where elderly reported on better results. No correlation could be found between monosyllabic speech perception in quiet and quality of life. However, in older subjects with better self-esteem speech understanding in quiet was better. Similar results have been found by Moberly

Table 5 Impact of Pre-operative Depressive Symptoms on Pre- and Post-operative Cognitive Performance, Quality of Life and Speech Perception According to Age

	Pre-operative Performance				Post-operative Performance			
	MA		OA		MA		OA	
	τ	p	τ	p	τ	p	τ	p
Cognitive subtests								
M3	0.31	0.03 *	-0.23	0.07	0.22	0.14	-0.03	0.79
Recall	0.08	0.57	0.003	0.98	0.04	0.76	-0.02	0.84
Delayed Recall	0.18	0.22	-0.04	0.75	0.25	0.09	-0.03	0.78
0-back	0.31	0.03 *	-0.08	0.55	0.11	0.43	-0.14	0.28
2-back	0.49	0.001 *	0.11	0.4	0.22	0.13	0.12	0.36
cFlanker	0.34	0.02 *	-0.09	0.47	0.15	0.32	-0.01	0.97
iFlanker	0.30	0.03 *	0.01	0.94	0.34	0.02 *	0.03	0.78
OSPAN	0.37	0.01 *	0.17	0.17	0.32	0.03 *	0.05	0.67
TMT A	0.39	0.01 *	0.23	0.07	0.32	0.03 *	0.29	0.02*
TMT B	0.32	0.03 *	0.13	0.32	0.17	0.25	0.18	0.15
Verbal fluency	0.01	0.96	-0.07	0.56	-0.03	0.83	0.14	0.27
Nijmegen Cochlear Implant Questionnaire								
Total score	-0.56	0.0002 *	-0.29	0.02 *	-0.09	0.52	-0.21	0.11
Physical	-0.61	0.00005 *	-0.31	0.01 *	-0.15	0.32	-0.26	0.047 *
Social	-0.28	0.06	-0.32	0.01 *	0.07	0.64	-0.16	0.2
Basic sound perception	-0.54	0.0003 *	-0.32	0.01 *	-0.04	0.8	-0.22	0.09
Advanced sound perception	-0.53	0.0004 *	-0.18	0.16	-0.18	0.24	-0.21	0.1
Speech production	-0.34	0.02 *	-0.1	0.44	-0.08	0.57	-0.14	0.26
Self-esteem	-0.49	0.001 *	-0.13	0.3	-0.15	0.31	-0.05	0.67
Activity limitations	-0.21	0.17	-0.34	0.01 *	-0.02	0.91	-0.16	0.21
Speech perception in quiet	-0.13	0.38	0.10	0.44	0.01	0.98	0.23	0.09

Notes: MA indicates middle-aged participants aged between 50 and 64 years and OA includes an age of ≥ 65 years. *Means significant ($p < 0.05$).

in 25 long-time CI users aged 50–83. Social interaction was correlated to an audio-visual task and to the recognition of complex sentences, but not to sentences in noise or to single words in quiet.⁶¹

However, as already stated by others improvement of quality of life after hearing rehabilitation is difficult to assess.^{62,63} Commonly used questionnaires that measure general quality of life such as the WHOQOL-BREF are time consuming and may not be suitable, because they do not encounter the impairment caused by hearing impairment⁶⁴ and health-related questionnaires as the Nijmegen Cochlear Implant Questionnaire that is widely used in hearing-impaired subjects do not sufficiently include daily activities.^{31,45,62,63,65} Therefore, both subjective self-assessments show only limited correlations to objective measurements.⁶⁶

This has also been found in a meta-analysis including 14 publications and covering 679 patients. Despite large post-operative improvements in quality of life only a low correlation could be observed between hearing in quiet or in noise and the Nijmegen Cochlear Implant Questionnaire, the

Hearing Handicap Inventory or the Abbreviated Profile of Hearing Aid Benefit.⁶³ Polku who investigated the effect of self-perceived hearing loss on quality of life measured by the WHOQOL-BREF observed that audiometrically assessed speech perception did not correlate to any QoL subdomain, whereas self-perceived impairment correlated to the total score and all subdomains (physical, psychological, environmental and social).⁶⁷ Sorrentino analyzed 69 CI users divided into 2 groups according to age (one older group of 25 individuals and a younger one of 19 individuals) as well as 25 normal hearing controls ≥ 65 with a cognitive screening test (Mini-Mental Status Examination) and the Glasgow Benefit Inventory commonly used to evaluate post-operative outcome in Otorhinolaryngology. Total QoL and the subdomain of physical health was related to speech perception, but only in the elderly and not in the younger aged group.⁶⁸

Concerning cognition, our findings demonstrate that neurocognitive functions significantly improve 12 months after cochlear implantation in middle-aged as well as in older adults with severe to profound hearing loss. But

whereas prior to surgery OA performed worse in working memory, inhibition, processing speed and mental flexibility, only inhibition significantly differed between the two age groups post-operatively. This is in line with data published by Salthouse showing that the ability to inhibit stimuli decreases with age.⁶⁹

Furthermore, Moberly found in a study on 30 hearing-impaired subjects with a mean age of 68.8 that the response time for inhibition assessed with a Stroop task was predictive for speech comprehension in noise.⁷⁰ This could also be observed in our study. The post-operative monosyllabic speech perception 12 months after implantation could be predicted pre-operatively by the inhibitory Flanker task in the older, but not in the middle-aged group.

Including age, hearing, gender and education as possible confounders, it was shown that hearing is a predictor for attention in both age groups and for OA with regard to inhibition. Cognitively challenging tasks such as the OSPAN, but also the Recall and the TMT are predominantly determined by education. Looking at the two age groups, age had a high impact on post-operative neuro-cognitive performance mainly on the 2-back, the compatible Flanker and on the Recall in older subjects in contrast to subjects <65 years of age.

Besides, a relationship has been found between health quality of life and some cognitive subtests: in MA advanced sound perception correlated to the 2-back task and improvement in working memory to a post-operative improvement of social interactions in the total group ($p=0.04$). Duration of hearing loss was associated with the total NCIQ score. Subjects with a shorter history of hearing impairment suffered more. Moberly described a negative correlation between patient's age and the sub-domain advanced sound perception and between a combination of patient's age at the time of implantation, duration of CI use, duration of hearing loss and the total well-being score.⁶¹

Due to the close connectivity between the auditory cortex and the limbic system, reduced peripheral input does not only lead to deactivation in central auditory structures but also to increased activation of cognitive control networks and to dysregulation of the limbic system on the neural level. On the behavioral level, social isolation may lead to depression as well as to cognitive decline in hearing impaired and mental disorders may be a mediator between hearing loss and cognitive decline.^{29,71}

CI candidates exhibit a wide range of psychological disorders as shown in a study by Brüggemann, who

described affective or somatoform disorders in 81% of adult CI candidates in contrast to 32% in the general population.⁷² In a review including 66 studies, Besser and colleagues also reported that hearing-impaired subjects are prone to suffer from multiple psychological disorders such as anxiety, an increased suicidal risk and social isolation.⁷³ Along with that Brewster observed in 3075 elderly aged 70–79 years that the risk to suffer from severe depression assessed by the CES-D scale (Centre for Epidemiologic Study Depression) increased 1.85-fold in hearing-impaired compared to normal hearing counterparts.⁷⁴ The use of hearing devices had no influence on mood. However, hearing evaluation was based on patient's report and not on detailed audiometric evaluation, and duration and frequency of hearing aid use were not encountered.⁷⁴

Dawes studied the association of cognitive performance and hearing status in hearing aid users as well as the relationship to social isolation and/or depression by structural equation modelling on a subsample of the UK Biobank data set ($n=164,770$) of UK adults aged 40–69.⁷⁵ Although a positive relationship has been detected between hearing aid use and cognition as well general health, there was no hint that this effect was mediated by social and psychological factors. Authors claim that hearing restoration might increase self-efficacy which is associated with better performance on challenging tasks. However, as subjects with higher cognitive scores might have searched more often for hearing devices a bias cannot be ruled out and longitudinal studies are missing.

In our study depression score was assessed only pre-operatively. Thereby the influence of hearing rehabilitation via cochlear implantation on depression could not be answered by this investigation. However, subjects who suffered from their hearing loss for a longer period of time showed less signs of depression in the younger aged group. Although severe depression has been excluded in our study population half of the subjects suffered from mild or moderate depressive symptoms. This is in line with a recent study by Tretbar in which only 22 out of 52 hearing impaired were without any psychiatric disorder, whereas 30 had been currently or previously treated, in 19 subjects due to symptoms of depression.⁷⁶

Knopke also studied psychosocial factors that might have an impact on quality of life such as the level of anxiety, stress and depression in 62 cochlear implant recipients aged 70–80 and in 24 CI users over 80. Whereas in the younger group NCIQ was mainly predicted by depressive symptoms,

anxiety was a positive predictor only for the octogenarians.⁷⁷ Age-related differences have also been described in a study of 50,398 Norwegians aged 20–101. Younger adults 20–44 and middle-aged adults of 45–64 perceived a higher psychosocial burden in depression, self-esteem and anxiety in case of hearing loss. In middle-aged the level of depressive symptoms increased by 0.1 SD for every 10dB hearing loss in contrast to elderly who showed only a slight association (0.01 SD increase for 10dB hearing loss).¹⁵

In our study, total depression score did not differ between the two age groups, but the impact of depression on cognitive performance and on well-being. In general, quality of life was poorer for patients with a higher level of depression. But whereas MA were significantly more affected by depression and had a higher improvement in quality of life if depression was higher, this association could not be observed in the elderly in the same way. It might be speculated that middle-aged hearing impaired who experience severe limitations in their daily activities at home or at work are more vulnerable to depressive symptoms than subjects aged ≥ 65 who are no longer part of the workforce and who might have accepted hearing loss as a part of normal aging.⁷¹ However, social support can moderate the relationship between hearing loss and depression in later life.⁷⁸ Additionally, the presence of depressive symptoms did not affect cognitive performance in the older group but in subjects <65 years of age.

A potential bias of the study might be due to the study design which evaluated the outcome mainly on questionnaires and behavioral investigation rather than on objective measures. At the moment normative scores or cut-off data for appropriate test-instruments for hearing impaired are still missing.^{79,80} Another limitation is that mental health assessed by the GDS-15 has only been done in a small sample size of 53 subjects and only prior to implantation. Other comorbidities such as anxiety were not studied.

To sum up cochlear implantation shows benefits in speech understanding and quality of life regardless of age. Neurocognitive functions which differ pre-operatively according to age also improve post-operatively. Subjects with worse pre-operative performance in cognitive and mental health show the greatest improvement. Quality of life is predicted by an interplay of various patients' characteristics such as age, but also by duration of hearing loss and cognitive as well as psychosocial variables.

Up to now, there is still limited knowledge about the benefit of hearing rehabilitation in severe cognitive or

depressive impaired older subjects.^{75,81,82} Further studies are required to figure out whether cochlear implantation might even improve quality of life in these patients as well as to identify the degree of impairment, which should not be exceeded to still achieve a benefit by cochlear implantation. To assess post-operative outcome, comorbidities should be taken into consideration as the impact of covariables strongly differs according to age.

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Author Contributions

CV, JPT, SD made substantial contributions to conception and design, LG, CV, IH to the acquisition of data, analysis and interpretation of data, CV, LG and JPT took part in drafting the article, IH and SD in revising it critically for important intellectual content; all authors gave final approval of the version to be published and agree to be accountable for all aspects of the work.

Disclosure

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