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# Dual innervation of free gracilis muscle for facial reanimation: What we know so far

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## Q4 KEYWORDS

Facial reanimation;  
Paralysis;  
Plastic surgery;  
Gracilis;  
Masseteric nerve;  
Graft;  
Dual innervation

## Abstract

**Background:** In the last decade, some institutions have begun combining the CFNG and masseteric nerve to provide dual innervation to the gracilis muscle for dynamic facial reanimation in facial paralysis patients. We reviewed the various ways that these two nerves have been coapted to provide dual innervation, and summarized the functional outcome for these methods.

**Methods:** A search of the Ovid EMBASE, MEDLINE, Cochrane, and Scopus databases was performed from 1946 to May 2019 for dual innervation of gracilis muscle using CFNG plus masseteric nerve for facial reanimation.

**Results:** A total of 184 articles were identified in the initial search, of which seven met our inclusion criteria. Three additional abstracts with 43 patients were identified but the level of details was not sufficient to include the results in the analysis. A total of 57 patients were reviewed (mean age of 42.1 years (6-79 years)). The majority of dual innervation procedures were performed using the ipsilateral masseteric nerve sutured end-to-end to the obturator nerve, and an additional CFNG connected end-to-side to the obturator nerve. In the 26 patients with Terzis scores available, there were no differences between masseteric nerve coapted end-to-end and CFNG as end-to-side to the obturator, or the reverse coaptation. All but two patients achieved function of the gracilis activated by the masseteric nerve within 2-5 months.

**Conclusions:** This review shows that dual innervation of the gracilis is safe; and in some cases, does appear to provide early onset gracilis activation as well as an eventual spontaneous smile.

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## 1 Introduction

2 In 1976, Harii et al. pioneered the use of a free functional  
3 gracilis muscle transfer innervated by the deep temporal  
4 nerve for facial reanimation.<sup>1</sup> At many institutions includ-  
5 ing ours, the gracilis is now the muscle of choice for reani-  
6 mation. Although CFNG provides optimal spontaneity, many  
7 surgeons continue to use the masseteric nerve due to the  
8 potential for many more axons to reach the muscle. Several  
9 authors have been using more than one nerve to provide  
10 innervation to the gracilis and connecting these nerves in  
11 various ways.

12 Early use of a trigeminal nerve branch as the sole power  
13 to the gracilis provided some symmetry when the patient  
14 would activate the gracilis by biting down, but it did not al-  
15 low for a predictable spontaneous smile.<sup>2,3</sup> In an effort to  
16 achieve spontaneous smile, several group have graft nerve  
17 from the unaffected facial nerve to the affected facial  
18 nerve or denervated muscles, referred to as a “cross face  
19 nerve graft” (CFNG).<sup>4-6</sup> In the late 1970s, O’Brien started  
20 using a two-staged technique: the first stage establishing  
21 the CFNG using a sural nerve sutured to the proximal buc-  
22 cal branches of the unaffected facial nerve, and the second  
23 stage performing a gracilis muscle transfer with coaptation  
24 to the already established CFNG.<sup>5,7</sup> He reported good to ex-  
25 cellent restoration of spontaneous smile in 51% of patients.<sup>7</sup>  
26 One of the drawbacks is the long distance required for the  
27 nerve to regrow, resulting in less nerve signals.<sup>12</sup> Combined  
28 with the multiple coaptations required, this could increase  
29 the risk of failed reinnervation.<sup>7,8</sup>

30 Recent literature have reported using a combination of  
31 both a CFNG and the masseteric nerve for innervation of a  
32 free functional gracilis transfer.<sup>9-15</sup> Theoretically, the fact  
33 that the masseteric nerve has a large number of axons, re-  
34 sults in faster recovery, stronger contraction, and a more  
35 symmetrical smile.<sup>3,16,17</sup> The CFNG contributes spontaneity  
36 that is hard to achieve consistently with masseteric nerve  
37 innervation alone. While conceptually appealing, it is not  
38 clear whether this method is comparable to the more sim-  
39 ple single innervation method, and how best to achieve this  
40 combined input.

41 We reviewed published surgical series utilizing a combi-  
42 nation of the masseteric nerve and the CFNG to re-innervate  
43 a transferred free functional gracilis muscle for facial re-  
44 animation. We were specifically interested in the descrip-  
45 tions of technique and associated clinical outcomes cur-  
46 rently available in the literature.

## 47 Methods

### 48 Literature search strategy

49 A comprehensive search was conducted across several  
50 databases including the Ovid EMBASE, MEDLINE, Cochrane,  
51 and Scopus databases for studies published through May  
52 2019. The search strategy was designed and conducted by  
53 a librarian with specialized training in literature retrieval.  
54 The search strategy was limited to human studies only. Key-  
55 words were used to search for cross facial nerve graft plus  
56 masseteric transfer. The actual terms used and how they  
57 are combined are shown in the search strategy (Appendix,

Supplemental Digital Content 1). In addition, because the  
58 search did not include abstracts presented at meetings,  
59 we specifically searched for published abstracts from rele-  
60 vant meetings including American Society of Plastic Surgery,  
61 American Society of Peripheral Nerve, American Society of  
62 Reconstructive Microsurgery, and Facial Nerve Symposium.  
63

### Inclusion and exclusion criteria

64  
65 Studies were included if they (1) reported data on patient  
66 demographics, outcomes, complications, and patient satis-  
67 faction and (2) were written in or translated into the English  
68 language. All age groups and sample sizes were included.

69 Studies were excluded if they were: (1) review papers,  
70 (2) pre-clinical studies, (3) technical notes, (4) animal stud-  
71 ies, (5) used only single innervation techniques, (6) if there  
72 was no free muscle flap used, or (7) if they used a free mus-  
73 cle other than the gracilis muscle.

### Selection of articles and data extraction

74  
75 Two authors (T.B. and J.M.) independently screened the ar-  
76 ticles through review of article titles and abstracts. Dupli-  
77 cates were then eliminated and an independent full text  
78 review of the remaining potentially relevant studies was  
79 performed using the exclusion and inclusion criteria. The  
80 extracted data included: year of publication, study design,  
81 total number of patients and number of procedures, type of  
82 technique, surgical outcomes, complications, and patient  
83 satisfaction. Disagreement between the reviewers was re-  
84 solved by discussion and consensus by a third independent  
85 reviewer (K.V.). This study complied with the guidelines out-  
86 lined in the Preferred Reporting Items for Systematic re-  
87 views and Meta-analyses (PRISMA).<sup>18</sup>

## 88 Results

### 89 Literature search strategy

90 **Figure 1** shows the PRISMA flow diagram of the performed  
91 search strategy. A total of 184 titles of potentially relevant  
92 publications were identified from the initial search strategy.  
93 After excluding 2 duplicates, 182 abstracts were screened,  
94 and 175 articles were excluded, with the most common rea-  
95 sons being the absence of free muscle transfer or only sin-  
96 gle innervation of the free muscle flap (see **Figure 1**, PRISMA  
97 flow diagram).

98 Full texts of 7 articles were reviewed and included in  
99 the final analysis (**Figure 1** and **Table 1**). All the articles in-  
100 cluded were published between 2012 and 2019. All studies  
101 were case series. The indication for all patients was long  
102 standing facial paralysis (see **Table 2**, which demonstrates  
103 the characteristics of included studies individually). In ad-  
104 dition, **Table 3** contains the included abstracts. These have  
105 been separated due to very limited information available.



## PRISMA 2019 Flow Diagram

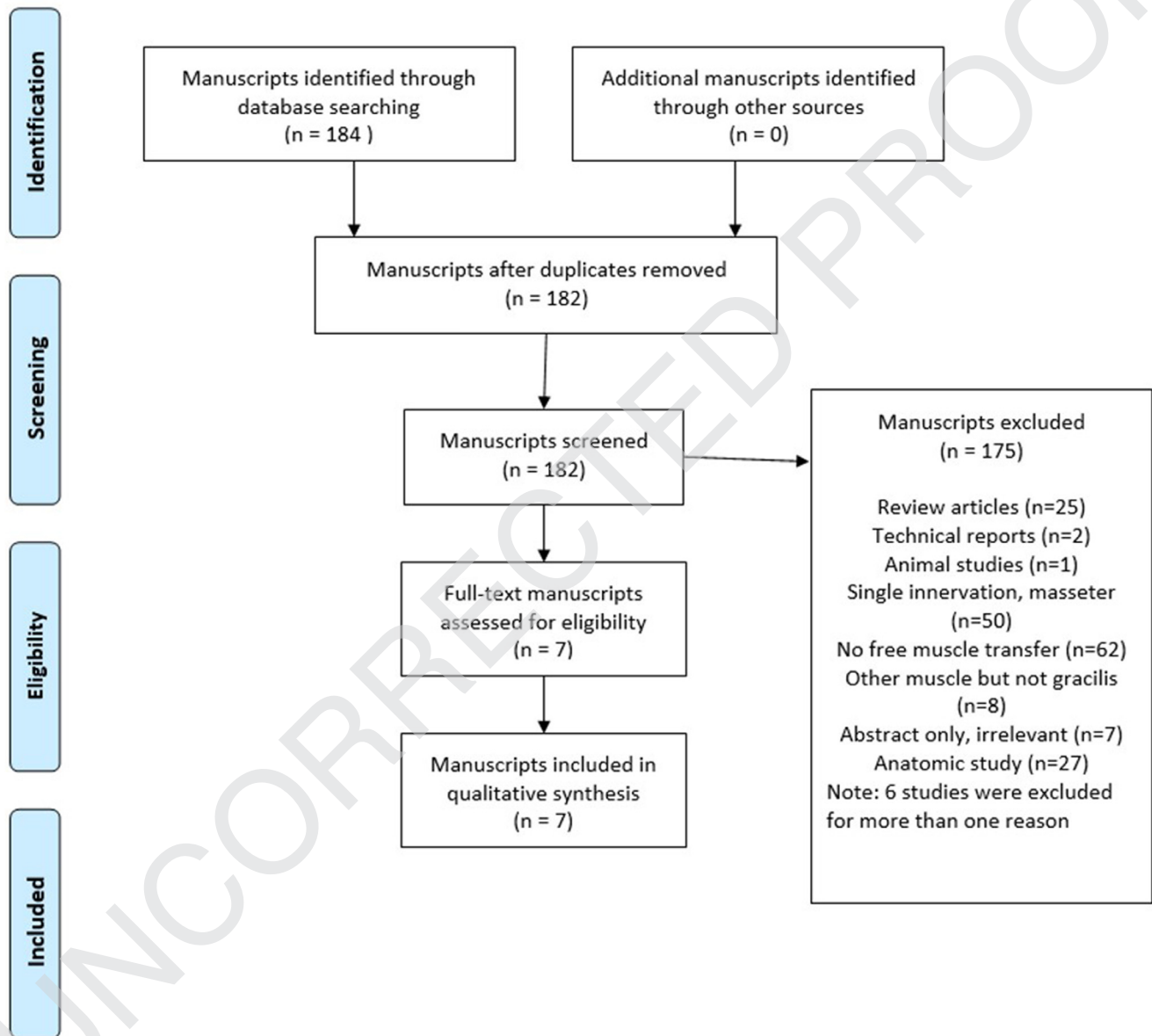


Figure 1 PRISMA flow diagram of the performed search strategy.

106 **Methods of dual innervation**

107 The majority of dual innervation procedures were per- 115  
 108 formed using the ipsilateral masseteric nerve coapted 116  
 109 end-to-end to the obturator nerve, with the CFNG sutured 117  
 110 end-to-side to the obturator nerve. All CFNG were sutured 118  
 111 end to end to the contralateral donor facial nerve (42 patients, 5 119  
 112 papers), see Figure 2. Cardenas-Mejia et al.<sup>13</sup> also discussed 120  
 113 performing the reverse innervation, with the CFNG as an 121  
 114 end-to-end coaptation to the obturator and the masseteric 122  
 123  
 124

nerve as an end-to-side coaptation to the obturator ( $n = 9$ ), 115  
 see Figure 3. Cardenas-Mejia et al.<sup>13</sup> were the only authors 116  
 to perform a two-stage procedure; all other authors per- 117  
 formed a single stage operation.<sup>10-12,14,15,19</sup> Cardenas-Mejia 118  
 et al. performed the masseteric nerve end to side coapta- 119  
 tion 1 cm from the gracilis muscle hilum. Sforza indicated 120  
 opening an epineural window for the end to side anastomosis.<sup>14</sup> 121  
 The other authors reviewed did not specify the exact 122  
 location or method of end to side anastomosis. Uehara and 123  
 Shimizu<sup>15</sup> discussed performing a procedure where they su- 124

**Table 1** Characteristics of studies describing free gracilis muscle transfer with dual innervation from CFNG and masseteric nerve for facial reanimation.

Method	CFNG as end-to-side coaptation and masseteric nerve as end-to-end	CFNG (1st stage) as end-to-end coaptation and masseteric nerve as end-to-side	End-to-end coaptations for both nerves using proximal and distal obturator
Number of papers	5 (Biglioli, Bianchi, Boahene, Sforza, Oh <sup>d</sup> )	1 (Cardenas-Mejia)	1 (Uehara)
Number of patients	42	9	6
Age (mean, range)(years)	39.6 (6-75)	37.6 (13-60)	56.7 (37-79)
Number of stages	1	2	1
Follow up (mean)(months)	Unclear in 1 paper; 18; 4 and 17	26.7	> 12
Number of patients achieving gracilis movement by the masseter	40 of 42 (95%)	9 of 9 (100%)	6 of 6 (100%)
Time to movement of transferred gracilis by masseter (average in months)	3.9 <sup>a</sup>	3.2	5.1
Was spontaneous smile achieved?	26 of 30 (86.6%) <sup>b</sup>	Not discussed	6 of 6 (100%)
Time to spontaneous smile (months)	Only discussed in one paper with mean 7.2 (6-8.8)	Not discussed	9.5
Terzis grading scale	Moderate (1/17, 5.8%), Good (9/17, 53%), Excellent (7/17, 41%) <sup>c</sup>	Moderate (1/9, 11%), Good (4/9, 44%), and Excellent (4/9, 44%)	Not used
Other grading scale used	Facial asymmetry index, and optoelectronic motion analysis 3D motion analyzer	N/A	Ratio of distance from angle of mouth when smiling compared to contralateral normal side

<sup>a</sup> With one paper where no evaluation was performed prior to 10 months, not included in the average.

<sup>b</sup> One paper with 5 patients did not discuss spontaneous smile.

<sup>c</sup> Two papers with 18 patients did not use Terzis score.

<sup>d</sup> For Suk et al., no average numbers are included in the table as they have 7 patients with the dual innervation and 3 patients with masseter only innervation. There was no breakdown of the data between these patients, and therefore we excluded their data in the average calculations such as follow-up period and time to movement of transferred gracilis.

125 tured an intramuscular motor branch of the gracilis to the  
126 ipsilateral masseteric nerve end-to-end, while the obtura-  
127 tor nerve was also sutured end-to-end to the CFNG ( $n = 6$ ),  
128 see Figure 4. The reviewed abstracts (Table 3) did not have  
129 details regarding the dual innervation method.

### 130 Outcome of clinical studies

131 Table 1 summarizes the general characteristics of the in-  
132 cluded studies including procedures, number of patients,  
133 follow-up, complications and outcomes. 57 patients (mean  
134 age of 42.1 years, range 6-79 years) underwent procedures  
135 with no serious adverse events reported. Table 3 summa-  
136 rizes the outcome of the abstracts. 43 patients from three  
137 abstracts were included, with one in the pediatric popula-  
138 tion and two in the adult population. There were no adverse  
139 events reported.<sup>20-23</sup> The Terzis Facial Grading System was  
140 used to evaluate outcomes in 26 patients.<sup>10,11,13</sup> The vast  
141 majority of these achieved good-to-excellent results, with  
142 no differences between the different methods of dual in-  
143 nervation coaptation. Pediatric patients were included in  
144 three series,<sup>9,13,14</sup> but there was not a detailed breakdown  
145 of individual outcomes, preventing the assessment of the in-

fluence of age on surgical outcome. In the abstract, McNeely  
146 et al.<sup>22</sup> presented 9 pediatric patients with age ranging from  
147 5 to 15 years old. They reported voluntary movement af-  
148 ter 4 months, with 3 reporting spontaneous movement by 3  
149 months. Continued improvement in all patients was noted  
150 until 12 months. The exact method of innervation was not  
151 specified. Note that Oh et al.<sup>19</sup> report a 10 patient case se-  
152 ries with 7 patients who underwent dual innervation. Unfor-  
153 tunately, as they did not separate their results, we had to  
154 exclude their data from select calculations such as age and  
155 time-to-spontaneous smile.

156 Follow-up time ranged widely from 4 months<sup>12</sup> to 18  
157 months.<sup>11</sup> In the abstracts, only mean follow time was given,  
158 with McNeely et al. reporting the longest follow up at mean  
159 27.33 months, with Win et al. and Dusseldorp et al. report-  
160 ing a 12 month follow-up.<sup>21-23</sup> All studies reported what they  
161 felt was the time to gracilis innervation from the masse-  
162 teric nerve, by assessing how long it took before the gracilis  
163 would contract when the jaw was clenched. 55 of 57 (96%)  
164 patients were able to smile while clenching/biting down, all  
165 within 2-5 months. The two abstracts that reported time-to-  
166 movement and spontaneous smile were by Win and Kallir-  
167 roi,<sup>23</sup> with results consistent with the full studies, and by  
168 McNeely et al.,<sup>23</sup> with results discussed above. Some studies  
169

**Table 2** Detailed breakdown of included manuscripts.

	No. of patients	Age (median, range)	No. of stages	Nerve anastomosis method	How was spontaneous smile assessed?	Time to movement of transferred gracilis by masseter (average in months)	Months to spontaneous smile	Rehab protocol?	Other grading scale used	Terzis Outcome classification	Follow up period
Bianchi et al., 2014	13	28 (6-73)	Single	CFNG as end-to-side coaptation and masseteric nerve as end-to-end	smile independent of biting contraction (no further details); "emotional activation"	3 (2-4 months); 3.9 months in the 4 unilateral gracilis transplantation with double innervation	not discussed; note all achieved spontaneity (4/4 and 9/9); dual gracilis - more rapid & powerful contraction than gracilis w/ CFNG alone	physiotherapy, smile independent from biting contraction	-	Dual: 2 good, 2 excellent, CFNG + masseter: 1 moderate, 5 good, 3 excellent	Unclear
Biglioli et al., 2012	4	49 (46-53)	Single	CFNG as end-to-side coaptation and masseteric nerve as end-to-end	observation by family member first detection, then by talking to patients and watch comedic movie for 10 mins	3.8 (2-4.8 months)	7.2 (6-8.8 months), all achieved spontaneous smile	physiotherapist guided exercises in front of them without mirror	EMG, and electroneurography. EMG: demonstrated reinnervation by CFNG when smiling w/o clenching; also demonstrated reinnervation by masseteric nerve during teeth clenching Electroneurography: able to directly stimulate gracilis via CFNG; unable to directly stimulate via masseteric nerve due to artefact from direct muscle stimulation.	2 excellent, 2 good	18 months

(continued on next page)

**Table 2** (continued)

	No. of patients	Age (median, range)	No. of stages	Nerve anastomosis method	How was spontaneous smile assessed?	Time to movement of transferred gracilis by masseter (average in months)	Months to spontaneous smile	Rehab protocol?	Other grading scale used	Terzis Outcome classification	Follow up period
Boahene et al., 2018	5	41(23-64)	3 Single, two patients 2 stages	CFNG as end-to-side coaptation and masseteric nerve as end-to-end	not specifically discussed, 'standard video analysis'	By 4 months, not specifically discussed	Not discussed.	Not discussed	dynamic smile zone analysis; Gingival and dentition analysis, Facial asymmetry index (FAI), using Canfield Mirror imaging software	Not discussed	4 months
Sforza et al., 2015	13	41 (9-75)	Single	CFNG as end-to-side coaptation and masseteric nerve as end-to-end	Funny video	10 months evaluation (sd, 5-16 months); all patients able to smile by clenching (2 patients did not regain any movement)	Not discussed; 9/12 patients able to perform spontaneous smile at a detectable level.	Not discussed	Optoelectronic motion analysis 3D motion analyzer; 'activation ratio' compared to healthy side.	Not discussed	17 months (SD 3)
Cardenas-Mejia et al., 2015	9	38 (13-60)	Two stages	CFNG (1st stage) as end-to-end coaptation and masseteric nerve as end-to-side	Not discussed	Time to reinnervation = 8.78 weeks (8-12 weeks); visible movement = 12.89 weeks (12-15)	Not discussed	Not discussed.	EMG used to assess reinnervation & motor unit recruitment	Moderate in 1, good in 4, and excellent in 4.	26.7 months (12-42)

(continued on next page)

**Table 2** (continued)

	No. of patients	Age (median, range)	No. of stages	Nerve anastomosis method	How was spontaneous smile assessed?	Time to movement of transferred gracilis by masseter (average in months)	Months to spontaneous smile	Rehab protocol?	Other grading scale used	Terzis Outcome classification	Follow up period
Uehara et al., 2017	6	57(37-79)	Single	Intramuscular branch of gracilis end to end to masseteric nerve, obturator transferred to CFNG.	Monthly evaluation, ask to smile with clenching and without clenching	4.7 months	9.5 months (9-12 months) (unclear clenching was involved or not, as only discussed 'synchronous' movement of the angle of the mouth bilaterally.	Biofeedback using mirror, >3 times daily.	Ratio of distance from angle of mouth when smiling compared to contralateral normal side, EMG, CMAP of CFNG.	Not used.	>18 months



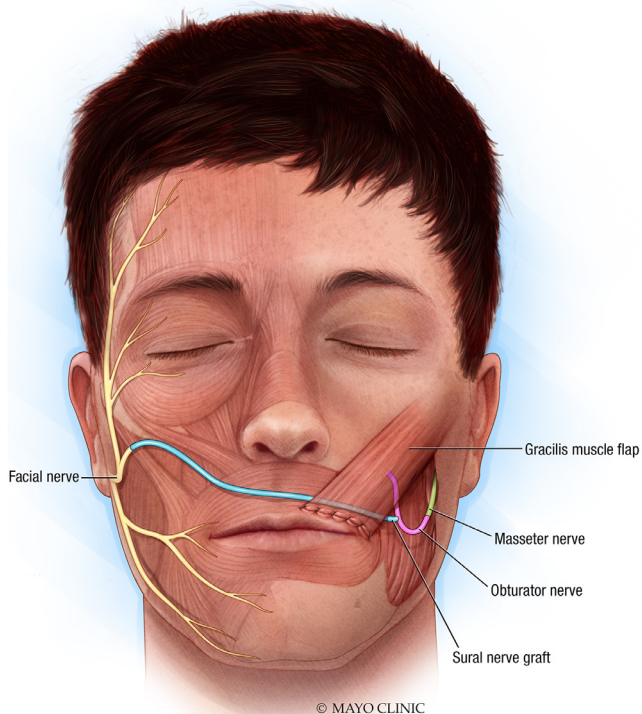
**Table 3** List of available abstracts from search of relevant meetings including American Society of Plastic Surgery, American Society of Peripheral Nerve, American Society of Reconstructive Microsurgery, and Facial Nerve Symposium.

	No. of patients	Age (median, range)	No. of stages	Nerve anastomosis method	How was spontaneous smile assessed?	Time to movement of transferred gracilis by masseter (average in months)	Months to spontaneous smile	Rehabs protocol?	Other grading scale used	Terzis Outcome classification	Follow up period
Win et al., 2014	8	35 (range 30-46)	2	Both CFNG and ipsilateral masseteric nerve	NA	3	10	Yes	EMG used	Used, but no results	12 months
McNeely et al., 2019	9	8.6 years (range: 5 to 15 years)	2	Both CFNG and ipsilateral masseteric nerve	NA	All patients demonstrated initiation of voluntary movement on the paralyzed side by 4 months, with three demonstrating spontaneous movement by 3 months. Improvements in excursion were noted to continue at 6 months, before stabilizing around 12 months.	Info in box to left	NA	House-Brackmann (HB) scores Eight patients had initial HB scores of VI and one patient had an initial HB score of V. Final HB ratings included five patients with a score of IV, three patients with a score of III and one patient with a score of II.	NA	Mean follow up was 27.33 months (SD 27.31)

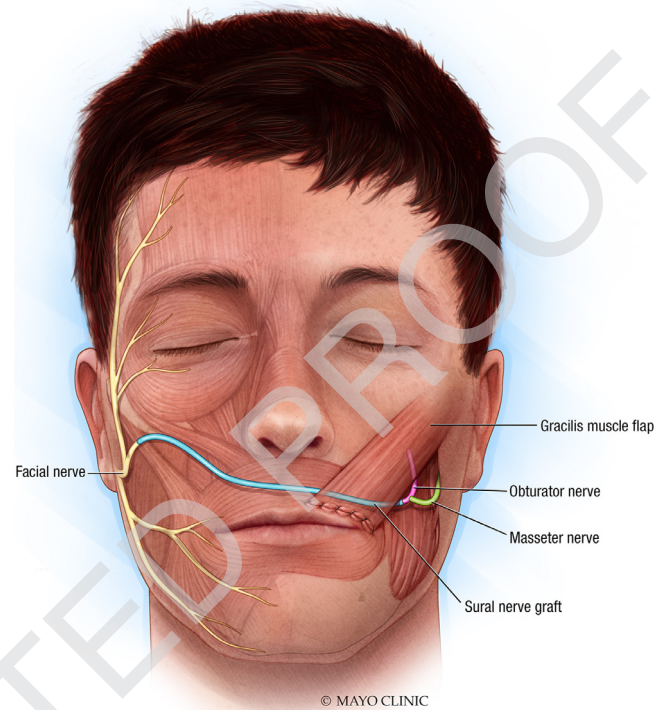
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**Table 3** (continued)

	No. of patients	Age (median, range)	No. of stages	Nerve anastomosis method	How was spontaneous smile assessed?	Time to movement of transferred gracilis by masseter (average in months)	Months to spontaneous smile	Rehabs protocol?	Other grading scale used	Terzis Outcome classification	Follow up period
Dusseldorp et al., 2018	26	NA	NA	Both CFNG and ipsilateral masseteric nerve	Validated humorous videos.	NA	NA	NA	eFACE and FaCE instrument A novel computer vision algorithm was employed to detect expression of joy during both voluntary and spontaneous smiling. eFACE and FaCE scale improvements were statistically significant. Results of both voluntary and spontaneous expression of joy in CFNG, NTM and dually innervated gracilis flaps will be presented.	NA	12



**Figure 2** Dual innervation using two end-to-end coaptations: CFNG with the obturator nerve, and the masseteric nerve with an intramuscular motor branch.



**Figure 3** Dual innervation using CFNG as an end-to-side coaptation to the obturator and the masseteric nerve as an end-to-end to the obturator nerve.

170 looked only at overall outcome, while others tried to differ- 196  
 171 entiate the strength of gracilis activation with biting down 197  
 172 versus a more emotional/spontaneous activation. Of the 36 198  
 173 patients where the presence of spontaneity was measured, 199  
 174 88% ( $n = 32$ ) were able to achieve a spontaneous smile. As 200  
 175 expected, the time to a spontaneous smile was delayed 201  
 176 compared to the time to gracilis activation with biting. Two 202  
 177 series (Biglioli et al. and Uehara et al.) assessed the time 203  
 178 to a spontaneous smile, reporting a mean of 7.2 months (6- 204  
 179 8.8) and 10 months (9-12), respectively.<sup>11,15</sup> McNeely et al. 205  
 180 reported spontaneous smile in 3 of 8 pediatric patients by 3 206  
 181 months.<sup>22</sup>

182 There were varied approaches to assessing whether a 207  
 183 smile was truly spontaneous or emotional in nature. Biglioli 208  
 184 et al., Bianchi et al., Sforza et al., and Dusseldorp et al. dis- 209  
 185 played funny videos to assess spontaneity.<sup>10,11,14,21</sup> Bianchi 210  
 186 et al. and Uehara et al. indicated that they specifically in- 211  
 187 structed the patient to not bite down during the assessment 212  
 188 for spontaneous smile.<sup>9,15</sup>

## 189 Neurophysiology and rehabilitation

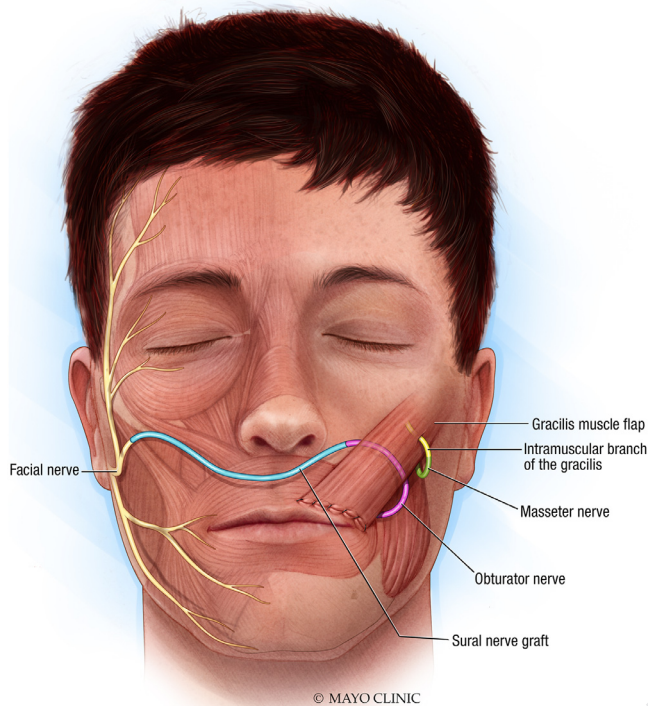
190 Three papers included electromyography (EMG) data. 213  
 191 Cardenas-Mejia et al. described a mean latency of 4.14 mil- 214  
 192 liseconds and a motor unit recruitment of 68.3% by one year 215  
 193 post-surgically, which they counted as a great outcome. 216  
 194 They did not specifically discuss how they stimulated the 217  
 195 muscle, however.<sup>13</sup> Biglioli et al. used EMG to verify the

CFNG function, by inserting an EMG coaxial needle elec- 196  
 trode into the gracilis muscle to assess motor units while 197  
 electrically stimulating the contralateral facial nerve.<sup>11</sup> Ue- 198  
 hara and Shimizu provided electrical stimulation of the con- 199  
 tralateral facial nerve at the tragus and upper lip and mea- 200  
 sured the motor potential amplitudes over the transferred 201  
 muscle on the affected side.<sup>15</sup> They reported compounded 202  
 motor action potentials ranging from 23 to 287 microvolts 203  
 in the 6 patients described. Interestingly, in the two cases 204  
 from their series with detailed descriptions of electrophys- 205  
 iologic testing, the gracilis flap was noted to be innervated 206  
 by the contralateral facial nerve and ipsilateral masseteric 207  
 nerve in one patient, and by only the masseteric nerve in 208  
 the other.

In addition to the surgical procedure, three series de- 210  
 scribed post-operative physical therapy, including reha- 211  
 bilitation protocols consisting of mirror biofeedback and 212  
 physiotherapist-guided exercises transitioning to smiling in- 213  
 dependent from biting.<sup>10,11,15</sup> In the abstracts, only Win 214  
 et al. reported use of EMG, but no further details were avail- 215  
 able.<sup>23</sup>

## 217 Discussion

218 Dual innervation of a free gracilis flap with a combination 219  
 of a CFNG and the ipsilateral masseteric nerve is a safe pro- 220  
 cedure with >95% of patients achieving some activation of 221  
 the gracilis by 2-5 months.



**Figure 4** Dual innervation using CFNG as an end-to-end coaptation to the obturator and the masseteric nerve as an end-to-side to the obturator nerve.

## 222 Comparison to single innervation methods

223 In single innervation studies, masseteric-innervated gracilis  
 224 muscles typically achieve a better excursion than CFNG-  
 225 innervated muscles, but lack a consistent spontaneity.<sup>2,3,17</sup>  
 226 Different methods of assessment were used, making ex-  
 227 cursión comparison difficult. Based on the timing to onset  
 228 of gracilis movement and the moderate to excellent re-  
 229 sults reported when the Terzis grading scale was applied,  
 230 dual innervation method achieved comparable results to se-  
 231 ries of gracilis muscle transfers using the masseteric nerve  
 232 alone.<sup>7,16,17,24</sup> Of those in whom the outcome of spontane-  
 233 ity was recorded, 88% (32-36) of patients were able to  
 234 achieve a spontaneous smile, similar to the spontaneity rate  
 235 reported in gracilis muscle transfers innervated by CFNG  
 236 alone.<sup>2,7,25,26</sup>

237 Time to gracilis reinnervation was reported by all au-  
 238 thors, ranging from 3.2 to 5.1 months. This is similar to  
 239 the 2-4 month time-to-activation of the gracilis muscle re-  
 240 ported with single innervation with the masseteric nerve.  
 241 The time to spontaneous smile as measured by Biglioli et al.  
 242 and Uehara et al. of 7.2 and 9.5 months is also consistent  
 243 with previously reported function after CFNG innervation of  
 244 about 1-2 year.<sup>2,3,7,27</sup> In the abstracts, Win et al.<sup>23</sup> reported  
 245 a time to gracilis reinnervation of 3 months, and time to  
 246 spontaneous smile of 10 months, but no further details were  
 247 given.

## Comparison between methods of dual innervation

248

249 It is too early in the description of dual innervated gracilis  
 250 transfers to be able to compare adequately the different  
 251 methods of innervation. Our review did not see any major  
 252 differences in outcome between the methods employed. We  
 253 have categorized the approaches into three groups as sum-  
 254 marized in Table 1 and Figures 2-4. The most common ap-  
 255 proach has been the use of the coaptation of the masse-  
 256 teric nerve end-to-end with the obturator with the CFNG as  
 257 an end-to-side coaptation to the obturator nerve.<sup>9-12</sup> Our  
 258 review also found an abstract-only article with an addi-  
 259 tional 8 patients using this approach, with good results.<sup>23</sup>  
 260 This would seem to put the masseteric nerve at a great  
 261 advantage, with any innervation from the CFNG acting as  
 262 a supplementary signal rather than providing the primary  
 263 nerve input. Cardenas-Mejia et al. described the reverse,  
 264 coapting the masseteric nerve end-to-side to the obturator  
 265 nerve, and the CFNG end-to-end to the obturator nerve, us-  
 266 ing two stages.<sup>13</sup> They reported a similar time to masseteric  
 267 re-innervation to other approaches. Uehara et al. found a  
 268 distal stump of the intramuscular motor branch of the ob-  
 269 turator nerve and used this as an end-to-end coaptation to  
 270 the masseteric nerve (Figure 4), therefore also permitting  
 271 end-to-end coaptation of the CFNG to the obturator nerve.<sup>15</sup>  
 272 With only one series describing this unique double end-to-  
 273 end innervation method, it is not clear yet whether this pro-  
 274 vides a better, similar, or worse outcome when compared to  
 275 other dual innervation techniques.

276 Snyder-Warwick et al.<sup>29</sup> studies the myelinated fiber  
 277 counts in their pediatric facial reanimation patients. The  
 278 downstream count in the CFNG at the second stage was  
 279 only 24% of the count at the facial nerve donor branch,  
 280 while the count from the masseteric nerve was 78%.<sup>29</sup> This  
 281 study confirmed the fact that the masseteric nerve provides  
 282 much stronger signal compared to the CFNG, which trans-  
 283 lated into a significant difference in the degree of move-  
 284 ment of the gracilis.<sup>29</sup> Several animal studies in rats also  
 285 looked into the differences between end-to-side vs. end-  
 286 to-end coaptations.<sup>30-32</sup> Both Liao et al. and Jaeger et al.  
 287 concluded that end-to-end coaptations of motor nerves re-  
 288 sulted in faster innervation and better muscle recovery fol-  
 289 lowing denervation compared to end-to-side innervation,  
 290 although the end-to-side method also provided reasonable  
 291 reinnervation potential.<sup>30,31</sup> In contrast, Viterbo et al.<sup>32</sup> did  
 292 not see any differences between the two methods. The work  
 293 by Isaac et al.<sup>33,34</sup> looking at the mechanism of end-to-side  
 294 coaptation, in addition to the clinical experience by Bar-  
 295 bour et al.<sup>35</sup> and Terzis et al.<sup>36</sup>, also supports end-to-side  
 296 coaptation as a viable method. Taking into account the find-  
 297 ings by Snyder-Warwick et al.<sup>29</sup> and Liao et al.,<sup>30</sup> Jaeger  
 298 et al.<sup>31</sup> and Viterbo et al.<sup>32</sup>, it is reasonable to conclude that  
 299 the masseter end-to-end method by Biglioli et al., Bianchi  
 300 et al., Sforza et al., and Oh et al.,<sup>9,11,14,19</sup> allows the mas-  
 301 seter nerve to provide the majority of the input to the gra-  
 302 cilis, while still allowing some signal from the CFNG through  
 303 end-to-side coaptation. The masseteric nerve as end-to-side  
 304 coaptation into the obturator as described by Cardenas-  
 305 Mejia et al.,<sup>13</sup> should allow more advantage to the CFNG  
 306 as this is coapted end-to-end, while still allowing strong  
 307 signal from the masseteric nerve. The abstract did not

308 detail the connection type, and was therefore not included  
309 here.

### 310 Single versus two-staged procedure

311 Both Biglioli et al. and Uehara et al. report using a single  
312 stage procedure, where the CFNG is placed at the same  
313 setting as the distal coaptation and the masseteric nerve  
314 transfer to the free gracilis flap.<sup>11,15</sup> Given that they per-  
315 formed the procedure in one stage, it seems unlikely that  
316 the longer CFNG would reach the contralateral side before  
317 the masseteric nerve. The animal experiment by Liao et al.  
318 further shows that end-to-side coaptation is slower and re-  
319 sults in less nerve input compared to end-to-end.<sup>30</sup> Because  
320 the masseteric nerve seems to innervate the gracilis first  
321 whether the procedure is performed in one or two stages,  
322 this may explain why the time-to-onset of gracilis activation  
323 was similar in all studies in our series.

### 324 Spontaneity of smile with dual innervation models

325 A common question is, how much of the patient's sub-  
326 sequent smile is due to the innervation originating from  
327 the masseteric nerve, and how much is from the CFNG?  
328 The most popular method of addressing spontaneity was  
329 by counseling the patients to smile without clenching their  
330 teeth, with the assumption that this would be considered a  
331 "spontaneous smile" and be attributed to the CFNG innerva-  
332 tion.<sup>10,11,14,15</sup> However, many patients with only masseteric  
333 nerve innervated gracilis transfers are eventually able to  
334 achieve a smile without biting down.<sup>37</sup> Manktelow et al.,<sup>37</sup>  
335 indicated in a patient questionnaire study that 69% of their  
336 patients with masseteric nerve innervated gracilis transfers  
337 learned to smile without biting down, after repeated prac-  
338 tice and training (average follow up of 4.7 years). In con-  
339 trast, Chuang et al., described a series of 22 patients with  
340 masseteric nerve innervated gracilis, and none of the pa-  
341 tients achieved a spontaneous smile using a "tickle test"  
342 (average follow up >2 years).<sup>2</sup> It is therefore difficult to  
343 know whether the patients in our review of the literature  
344 who were reported to have a spontaneous smile by "smiling  
345 without clenching" achieved this spontaneity based on the  
346 additional CFNG, or cortical plasticity.<sup>38,39</sup> Perhaps a better  
347 method of evaluating true spontaneity and even an emotive  
348 smile was done by Biglioli et al., Bianchi et al., Sforza  
349 et al. and Dusseldorp et al., when they measured gracilis  
350 activation after a funny video.<sup>9,11,14,21</sup> However, even this  
351 level of spontaneity has been reported with masseteric in-  
352 nervation alone.<sup>38</sup> In the pediatric patients reported by Mc-  
353 Neely et al.<sup>22</sup> patients appear to achieve spontaneous smile  
354 earlier, with 3 of 9 with spontaneous smile by 3 months.  
355 Whether the entire group of 9 patients achieved sponta-  
356 neous smile was not clear from the abstract. Future studies  
357 using dual innervation gracilis transfer models may benefit  
358 from EMG evaluations similar to those used by Uehara et al.  
359 and Biglioli et al., where the CFNG input is assessed more  
360 objectively through measurement of the CFNG compound  
361 muscle action potential (CMAP) and by stimulating the prox-  
362 imal CFNG while measuring the motor unit response in the  
363 gracilis muscle.<sup>11,15</sup>

The case series that were available in the literature did  
not use consistent methods for post-surgical evaluation,  
making direct comparisons difficult. Also, there are surgi-  
cal centers that may be performing dual innervation of the  
gracilis muscle who have not yet published, and for this rea-  
son we tried to review the available abstracts in the liter-  
ature as well. While these abstracts did provide some ad-  
ditional details on outcome, detailed evaluations were not  
available. Finally, it is possible that there has been publi-  
cation bias preventing publication of negative results from  
these or additional novel dual innervation models. This high-  
lights the need for further publications with standardized  
electrophysiological and clinical outcome measures.

### 377 Conclusions

378 Surgical centers have only started to incorporate dual in-  
379 nervated gracilis transfers for facial reanimation in the past  
380 6 years. It is still too early to know if one specific method  
381 of coaptation yields a better surgical outcome than others.  
382 Based on the available literature to date, we do know that  
383 the dual innervated free gracilis muscle utilizing the mas-  
384 seteric nerve and CFNG seems to be a safe procedure, with  
385 gracilis activation similar in timing to single innervation pro-  
386 cedures using just the masseteric nerve, and a substantial  
387 number of patients also achieving spontaneity.<sup>10,11,14,15,37</sup>

In order to better evaluate these procedures and deter-  
mine the optimal approach, future studies should be de-  
signed with standardized measures of spontaneity, including  
the incorporation of emotive stimuli (such as funny videos or  
"the tickle test") and electrophysiologic techniques.

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### 402 Declaration of Competing Interest

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## Supplementary materials

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