

TMR Integrative Medicine

Botulinum toxin type A combined with robot-assisted training for upper limb spasticity and motor function after stroke: a case report

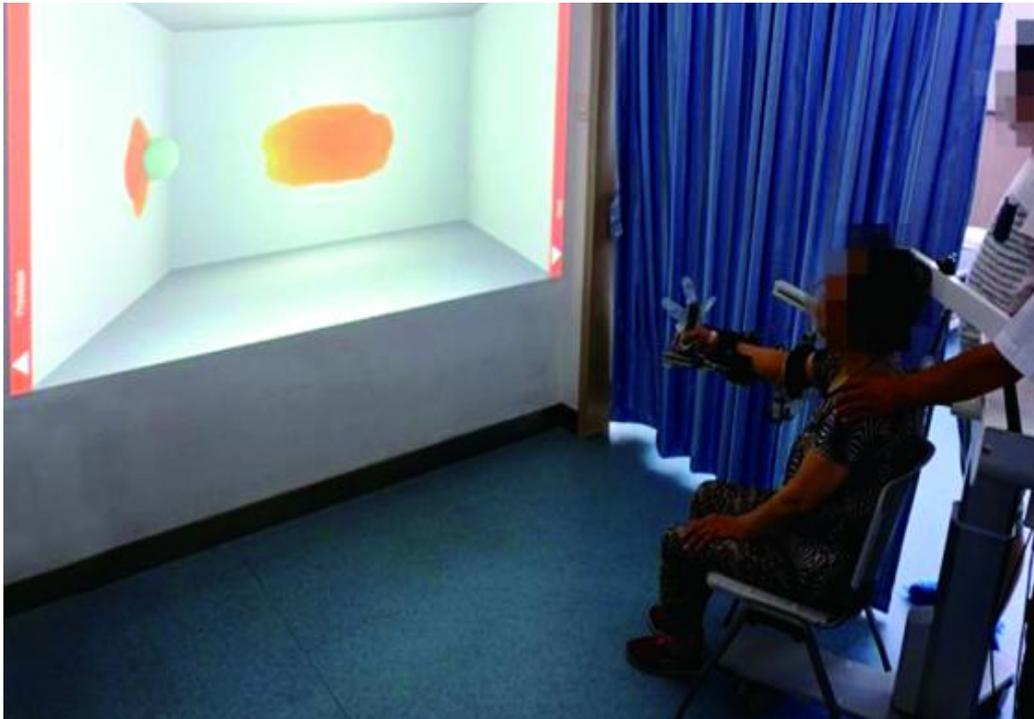
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Highlights

The study shows that Botulinum toxin type A in combination with robot-assisted training treatment is a viable training method for improving upper limb spasticity and motor function, which provides a new reference value for clinical therapy in patients with spastic stroke.



Abstract

This study is to report the clinical experience of Botulinum toxin type A in combination with robot-assisted training on upper limb spasticity and motor function in stroke patient through a case report. The patient underwent combined treatment with Botulinum toxin type A and robot-assisted training of the upper limb for 3 weeks. Evaluation was performed before and after combined treatment, and 3 months follow-up after discharge. The following outcomes were measured: spasticity by the modified Ashworth scale, pain by the numeric rating scale, motor function by the Fugl-Meyer assessment in upper limb, and activity of daily living by the modified barthel index. The combined Botulinum toxin type A and robot-assisted training treatment had an obvious improvement in upper limb spasticity, pain, motor function and activity of daily living. Botulinum toxin type A combined with robot-assisted training therapy is worthy of further application in patients with spastic stroke.

Keywords: Botulinum toxin type A, Robot, Spasticity, Motor function, Stroke

Abbreviations:

BTX-A, Botulinum toxin type A; 3D, 3-dimensional; ADL, activity of daily living; MAS, modified Ashworth scale; NRS, pain by the numeric rating scale; FMA-UL, Fugl-Meyer assessment in upper limb; MBI, modified barthel index.

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Competing interests:

The authors declare that there is no conflict of interest. Written informed consents were obtained from patient.

Author Contributions:

Min Zhang and Hong You designed the study protocol. Yong-Ping Li, Ming-Ming Wen collected the data. Min Zhang drafted the initial manuscript. All authors contributed to the revision of the manuscript and have read and approved the final version.

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Background

Hemiplegia spasticity is one of the most common disability manifestations of stroke patients, which is mainly manifested by increased muscle tension of upper and lower limbs on the affected side, accompanied by symptoms such as pain, muscular atrophy and limb motor dysfunction. It is easy to cause anxiety and depression of patients, which seriously affects the daily life and rehabilitation efficacy of patients [1].

Traditional rehabilitation training (such as Bobath neurodevelopmental techniques, functional mobility training, oral medications, etc.) is the main treatment to help restore the limbs function, while it is also time consuming and labor-intensive processes. Moreover, traditional rehabilitation training is slow-acting and more appropriate for treating chronic ailments, which may lead to miss the best time for treatment [2, 3]. At present, in order to improve the upper limb motor function and training enthusiasm of stroke patients,

besides relying on therapists to carry out some traditional rehabilitation training, robot-assisted training has been gradually applied to clinical practice, it uses computer virtual software to create a virtual space close to the life environment for patients, and trains stroke patients through sight, hearing and touch, so as to promote the recovery of their upper limb function [4-8].

Studies have shown that Botulinum toxin type A (BTX-A) therapy can improve upper limb spasticity, relieve pain and improve joint mobility after stroke, and at the same time, it takes effect faster and lasts longer than traditional rehabilitation training, and shortens the rehabilitation process [9-11]. However, the effect of robot-assisted training in combination with BTX-A therapy on upper limb spasticity and motor function after stroke is not clear. Thus, the purpose of this study is to report a case describing the effects of BTX-A in combination with robot-assisted training on upper limb spasticity and motor function in stroke patient.

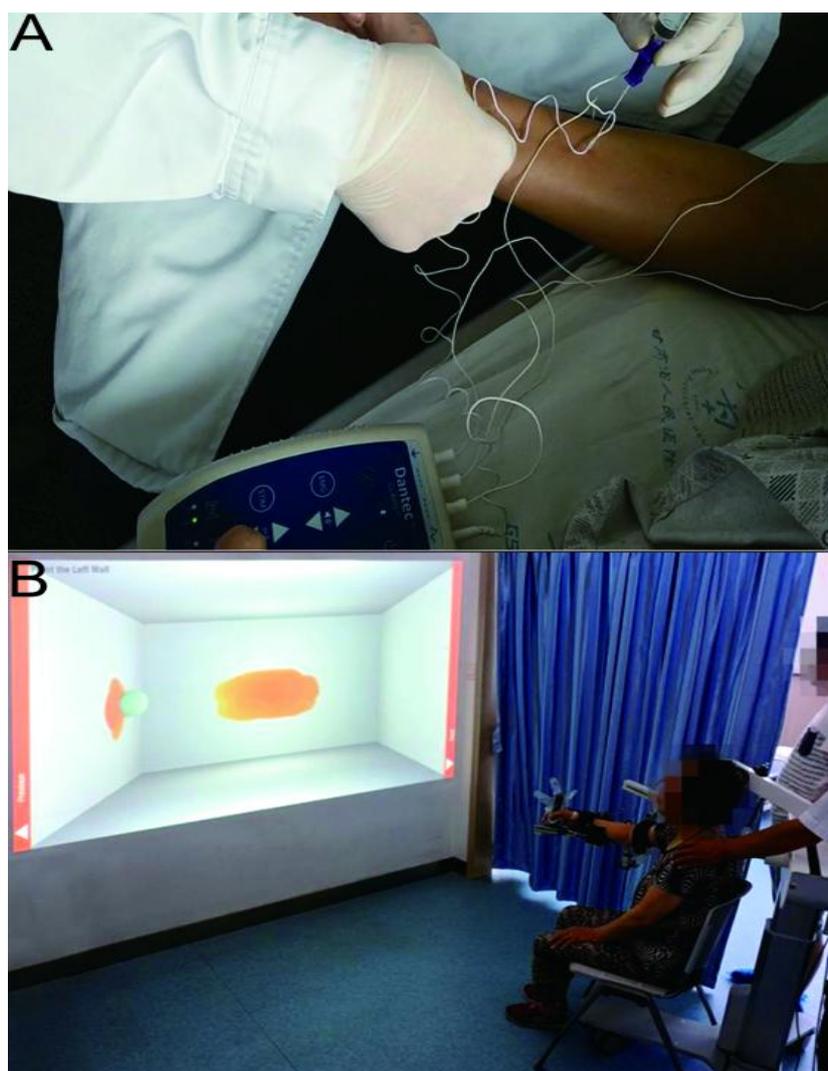


Figure 1 Patient received the combination of BTX-A (A) and robot-assisted training treatment (B)

Table 1 The effects of the combination of BTX-A and robot-assisted training treatment

	Pre-treatment	Post-treatment	Follow-up
MAS			
Biceps brachii	3	2	1
Brachioradialis	3	2	1
Pronator teres	2	1	1
Flexor digitorum superficialis	2	1	1
Flexor digitorum profundus	3	2	2
NRS	7	2	2
FMA-UL	40	51	52
MBI	52	76	78

MAS, modified Ashworth scale; NRS, pain by the numeric rating scale; FMA-UL, Fugl-Meyer assessment in upper limb; MBI, modified barthel index.

Case description

On May 3, 2018, a 53-year-middle aged man, went to the Sino-French Department of Neurological Rehabilitation of Gansu Provincial Hospital due to right hemiplegia and severe spasticity of the upper limb, presented with left basal ganglia hemorrhage from 3 months ago.

After admission, regular physical examination was conducted, the patient was consciousness but had speech deficient, with T: 36.3°C, P: 93 times/min, R: 19 times/min, BP: 158/98 mmHg; both pupils equal in size and diameter about 3 mm, sensitive to light reflexes, bilateral eye movements in all directions freely in place, no nystagmus or diplopia, soft neck without resistance, bilateral nasolabial sulcus symmetrical, tongue extended and centered, normal pharyngeal reflexes, right upper limb muscle strength was grade III, right lower limb muscle strength was grade IV, left upper and lower limb muscle strength was grade V, right upper limb muscle tone was increased, limb tendon reflexes were normal, and bilateral pathological signs were negative.

Cranial CT showed left basal ganglia hemorrhagic foci absorbed.

The patient had a history of hypertensive disease and cerebral hemorrhage, and took oral amlodipine for blood pressure control and mecobalamin for neuroprotection (regular dose).

The treatment of BTX-A (Institute of Biological products, Lanzhou, China) was injected in different upper limb muscles (Figure 1A). Detail as follows: Biceps brachii (100 unit), Brachioradialis (60 unit), Pronator teres (40 unit), Flexor digitorum superficialis (40 unit), Flexor digitorum profundus (60 unit). Robot-assisted training started the day after the BTX-A injection and continued for twice a day (30 minutes per time), 5 times a week for a total period of 3 weeks. The robot (Hocoma Inc, Switzerland. Figure 1B) is a 3-dimensional (3D), multi-joint rehabilitation training and evaluation system that can provide accurate gravity compensation, its training program is 3D

motion execution based on virtual reality, include movements of shoulder, elbow and wrist joints, and hand grasping, which are similar to real activity of daily living (ADL) action.

Assessment of the patients response to therapy was monitored by the use of the following assessment methods: the modified Ashworth scale (MAS) to measure spasticity grade [12, 13], Numerical Rating Scale (NRS) to measure pain intensity [14, 15], the Fugl-Meyer assessment in upper limb (FMA-UL) to measure motor function [16], and the modified barthel index [17] (MBI) to measure ADL. Evaluations were performed before the BTX-A injection combined with robot-assisted training, after combined treatment and follow-up data were taken 3 months after discharge, the results of evaluations showed that the combined therapy had greatly clinical improvement in the MAS, NRS, FMA-UL, MBI of this stroke patient (Table 1). In concrete terms, there was decreased in pain and spasticity, increased in motor function and ADL that were maintained up to three-month follow-up after discharge.

Discussion

In this case report, we observed the improvements of a post-stroke patient after 3 weeks of robot-assisted in combination with BTX-A therapy. In previous study, Botulinum toxin type A was found to be safe and efficacious for the treatment of poststroke limb spasticity [18–21]. In addition, it had reported the effectiveness of BTX-A on the upper limb pain [22–24]. These outcomes on spasticity and pain are similar to this case report. Robot-assisted training can promote the recovery of the upper limb motor function and ADL in this study. We considered that robot-assisted training system can provide upper limb support and increase sensory information input based on virtual reality, thus patient can easily carry out independent movement, and effectively improve the upper limb motor ability and ADL through active training, some similar studies have the same viewpoint [25, 26]. About the effects of robot-assisted training on

spasticity, the improvements may be due to counteracting the influence of gravity on the upper limb active training, reducing the requirements of the patients' active muscle movement function and physical strength, making the active muscle and the antagonistic muscle repeatedly balance training, inhibiting the excessive increase of muscle tension and the occurrence of abnormal movement mode [27]. However, further studies are necessary to demonstrate this theory for robot-assisted training. After 3 months of follow-up period, it was considered to have reached a platform period of upper limb function at discharge. As this is a case report, the study results are universal, but they can expand the number of cases and select the control group to further test the combined treatment method in clinical trials.

Conclusion

The combined BTX-A and robot-assisted training treatment had an obvious improvement in upper limb spasticity, pain, motor function and ADL. Therefore, the combination therapy is worthy of further application in patients with spastic stroke.

References

- Merriman C, Norman P, Barton J. Psychological correlates of PTSD symptoms following stroke. *Psychol Health Med* 2007, 12: 592–602.
- Dou W, Liu XY, Ma YN, et al. Effect of botulinum toxin type A combined with virtual training on upper limb functional recovery and analysis of the mechanism after stroke. *Chin J Trauma Disabil Med* 2019, 27: 84–86.
- Jiang Y, Xie RM, Cao HG, et al. Effect of type A botulinus toxin combined with targeted training on the wrist function and pain indexes in patients with stroke complicated by spasm. *Hainan Med J* 2016, 27: 1445–1447.
- Kwakkel G, Kollen BJ, Krebs HI. Effects of robot-assisted therapy on upper limb recovery after stroke: a systematic review. *Neurorehabil Neural Rep* 2007, 22: 111–121.
- Masiero S, Armani M, Rosati G. Upper-limb robot-assisted therapy in rehabilitation of acute stroke patients: Focused review and results of new randomized controlled trial. *J Rehabil Res Devel* 2011, 48: 355–366.
- Frisoli A, Procopio C, Chisari C, et al. Positive effects of robotic exoskeleton training of upper limb reaching movements after stroke. *J Neuroeng Rehabil* 2012, 9: 36.
- Masiero S, Armani M, Ferlini G, et al. Randomized trial of a robotic assistive device for the upper extremity during early inpatient stroke rehabilitation. *Neurorehabil Neural Rep* 2013, 28: 377–386.
- Klamroth-Marganska V, Blanco J, Campen K, et al. Three-dimensional, task-specific robot therapy of the arm after stroke: a multicentre, parallel-group randomised trial. *Lancet Neurol* 2014, 13: 159–166.
- Xu GQ, Lan Y, Zhao JL, et al. Dilution of botulinum toxin A in lidocaine vs. in normal saline for the treatment of upper limb spasticity in stroke patients: a randomized, comparative study. *Chin J Rehabil Med* 2015, 30: 237–241.
- Simpson DM, Gracies JM, Yablon SA, et al. Botulinum neurotoxin versus tizanidine in upper limb spasticity: a placebo-controlled study. *J Neurol Neurosurg Psychiatry* 2009, 80: 380–385.
- Yang YM, Liang Q, Wan XH, et al. Safety and efficacy of botulinum toxin type A made in China for treatment of post-stroke upper limb spasticity: a randomized double-blind controlled trial. *Chin J Neurol* 2018, 51: 355–363.
- Kong KH, Chua KS, Lee J. Symptomatic upper limb spasticity in patients with chronic stroke attending a rehabilitation clinic: frequency, clinical correlates and predictors. *J Rehabil Med* 2010, 42: 453–457.
- Lundström E, Smits A, Terént A, et al. Time-course and determinants of spasticity during the first six months following first-ever stroke. *J Rehabil Med* 2010, 42: 296–301.
- Donald DP, Francis MB, Stephen L, et al. A comparison of pain measurement characteristics of mechanical visual analogue and simple numerical rating scales. *Pain* 1994, 56: 217–226.
- Wood BM, Nicholas MK, Blyth F, et al. Assessing pain in older people with persistent pain: the NRS is valid but only provides part of the picture. *J Pain* 2010, 11: 1259–1266.
- Fugl-Meyer AR, Jääskö L, Leyman I, et al. The post stroke hemiplegic patient. 1. a method for evaluation of physical performance. *Scandinavian J Rehabil Med* 1975, 7: 13–31.
- Shah S, Vanclay F, Cooper B. Improving the sensitivity of the Barthel Index for stroke rehabilitation. *J Clin Epidemiol* 1989, 42: 703–709.
- Das TK, Park DM. Effect of treatment with botulinum toxin on spasticity. *Postgraduate Med J* 1989, 65: 208–210.
- Rosales RL, Chua-Yap AS. Evidence-based systematic review on the efficacy and safety of botulinum toxin-A therapy in post-stroke spasticity. *J Neural Transm* 2008, 115: 617–623.
- Rodgers H, Shaw L, Price C, et al. Study design and methods of the BoTULS trial: a randomised controlled trial to evaluate the clinical effect and cost effectiveness of treating upper limb spasticity due to stroke with botulinum toxin type A. *Trials* 2008, 9: 59.
- Elovic EP, Brashear A, Kaelin D, et al. Repeated

- treatments with botulinum toxin type A produce sustained decreases in the limitations associated with focal upper-limb poststroke spasticity for caregivers and patients. *Arch Phy Med Rehabil* 2008, 89: 799–806.
22. Lim JY, Koh JH, Paik NJ. Intramuscular botulinum toxin-A reduces hemiplegic shoulder pain: a randomized, double-blind, comparative study versus intraarticular triamcinolone acetonide. *Stroke* 2008, 39: 126–131.
 23. Yelnik AP, Colle FM, Bonan IV, et al. Treatment of shoulder pain in spastic hemiplegia by reducing spasticity of the subscapular muscle: a randomised, double blind, placebo controlled study of botulinum toxin A. *J Neurol Neurosurg Psychiatry* 2007, 78: 845–848.
 24. Bhakta BB, Cozens JA, Bamford JM, et al. Use of botulinum toxin in stroke patients with severe upper limb spasticity. *J Neurol Neurosurg Psychiatry* 1996, 61: 30–35.
 25. Liao WW, Wu CY, Hsieh YW, et al. Effects of robot-assisted upper limb rehabilitation on daily function and real-world arm activity in patients with chronic stroke: a randomized controlled trial. *Clin Rehabil* 2011, 26: 111–120.
 26. Norouzi-Gheidari N, Archambault PS, Fung J. Effects of robot-assisted therapy on stroke rehabilitation in upper limbs: Systematic review and meta-analysis of the literature. *J Rehabil Res Dev* 2012, 49: 479–496.
 27. Cauraugh JH, Summers JJ. Neural plasticity and bilateral movements: a rehabilitation approach for chronic stroke. *Prog Neurobiol* 2005, 75: 309–320.