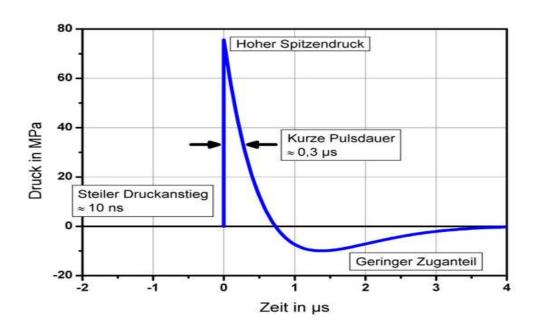
# Stoßwellentherapie bei Lymphödem

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# Stoßwelle - Physik





Quelle: www.digest-ev.de

Quelle: Wikipedia



## Wirkprinzipien

	Upregulation Factors	Downregulation Factors
Chondroprotective effect <sup>2,14,16,29,40</sup>	BMP-2, 3, 4, 7; IGF-1; TGFβ-1; VEGF; Wnt3; RUNX2; osteocalcin; alkaline phosphatase; osteopontin; FAK; ERK1/2; c-Fos; c-Jun; p38 MAPK; P2X7 receptor; S0X9; PDGF; b-FGF; FGF-2; Ras; substance P; prostaglandin E(2); Hsp70	DKK1, Wnt5a, calcitonin gene-related peptide miR-138
Neovascularization 25,43-45	VEGF, Flt1, Flt2, CD31, vWF, FGF, PIGF, KDR, PCNA	
Anti-inflammation 12,13,22,25,46	TGFβ-1, TLR3, eNOS, nNOS, IL-10, IL-6, IL-8, cyclophillin B, cyclophillin A, EGF-like domains 2, IFN-β1	sICAM, sVCAM, iNOS, IL-18, TNF $\alpha$ , NF-kB
Anti-apoptosis <sup>25,47</sup>	Bcl2, heme oxygenase (HO)-1, NAD(P)H quinone oxidoreductase-1	Bax, cleaved caspase 3, cleaved PARP, γ-H2AX, NOX1, NOX2, TUNEL activity
Tissue and nerve regeneration <sup>35,48-52</sup>	COL1A1, COL2A1, MMP2, MMP9, glycosaminoglycan, collagen type III, S100b, p75, c-Jun, GFAP, activating transcription factor 3 (ATF3), growth-associated phosphoprotein (GAP-43)	MMP-1, MMP-13, myelin marker P0

\*BMP = bone morphogenetic protein, IGF = insulin-like growth factor, TGF = transforming growth factor, VEGF = vascular endothelial growth factor, RUNX2 = runt-related transcription factor 2, FAK = focal adhesion kinase, ERK = extracellular signal-regulated kinase, MAPK = mitogen-activated protein kinase, PDGF = platelet-derived growth factor, FGF = fibroblast growth factor, Hsp = heat-shock protein, DKK = dickkopf-related protein, miR = microRNA, FIt = FMS-like tyrosine kinase, vWF = von Willebrand factor, PIGF = phosphatidylinositol-glycan biosynthesis class-F protein, KDR = kinase insert domain receptor, PCNA = proliferating cell nuclear antigen, TLR = Toll-like receptor, NOS = nitric oxide synthase, eNOS = endothelial NOS, nNOS = neuronal NOS, IL = interleukin, EGF = epidermal growth factor, IFN = interferon, sICAM = soluble intercellular adhesion molecule, sVCAM = soluble vascular cell adhesion molecule, iNOS = inducible NOS, TNF $\alpha$  = tumor necrosis factor alpha, NFkB = nuclear factor kappa B, Bcl = B-cell lymphoma, NAD = nicotinamide adenine dinucleotide, PARP = poly(ADP-ribose) polymerase, H2AX = H2A histone family member X, TUNEL = terminal deoxynucleotidyl transferase-mediated dUTP nick end labeling, COL1A1 = collagen type-1 alpha 1, COL2A1 = collagen type-2 alpha 1, MMP = matrix metalloproteinase, and GFAP = glial fibrillary acidic protein.

The Role of Extracorporeal Shockwave Treatment in Musculoskeletal Disorders Moya, Daniel MD<sup>1,a</sup>; Ramón, Silvia MD, PhD<sup>2</sup>; Schaden, Wolfgang MD<sup>3</sup>; Wang, Ching–Jen MD<sup>4</sup>; Guiloff, Leonardo MD<sup>5</sup>; Cheng, Jai–Hong MD<sup>4</sup> JBJS: February 7, 2018 – Volume 100 – Issue 3 – p 251–263



### ESWT Indikationen, ISMST The International Society for Medical Shockwave Treatment

46 Indikationen für Stoßwellentherapie

- **Approved standard indications**
- Common empirically-tested clinical uses
- **Exceptional indications expert indications**
- **Experimental Indications**

Quelle: www.shockwavetherapy.org



### ESWT Indikationen, ISMST The International Society for Medical Shockwave Treatment

#### 3. Exceptional indications - expert indications

- Musculoskeletal pathologies
- 3.1.1. Osteoarthritis
- 3.1.2. Dupuytren disease
- 3.1.3. Plantar fibromatosis (Ledderhose disease)
- 3.1.4. De Quervain disease
- 3.1.5. Trigger finger
- 3.2. Neurological pathologies
- 3.2.1. Spasticity
- 3.2.2. Polyneuropathy
- 3.2.3. Carpal Tunnel Syndrome
- 3.3. Urologic pathologies
- 3.3.1. Pelvic chronic pain syndrome (abacterial prostatitis)
- 3.3.2. Erectile dysfunction
- 3.3.3. Pevronie disease
- 3.4. Others
- 3.4.1. Lymphedema

Quelle: www.shockwavetherapy.org



## Experimentelle Untersuchungen

#### Extracorporeal shock wave therapy ameliorates secondary lymphedema by promoting lymphangiogenesis

Masayuki Kubo, PhD, Tao-Sheng Li, MD, PhD, Takahiro Kamota, MD, Mako Ohshima, PhD, Bungo Shirasawa, MD, PhD, and Kimikazu Hamano, MD, PhD, Ube, Yamaguchi, Japan

Objective: Although secondary lymphedema is a common complication after surgical and radiation therapy for cancer, the treatment options for lymphedema remain limited and largely ineffective. We thus studied the effect of extracorporeal shock wave therapy on promoting lymphangiogenesis and improving secondary lymphedema.

Methods: A rabbit ear model of lymphedema was created by disruption of lymphatic vessels. Two weeks after surgery, the lymphedematous ear was treated with or without low-energy shock waves (0.09 mJ/mm<sup>2</sup>, 200 shots), three times per

Results: Western blot analysis showed that the expression of vascular endothelial growth factor (VEGF)-C (1.23-fold, P < .05) and VEGF receptor 3 (VEGFR3; 1.53-fold, P < .05) was significantly increased in the ears treated with shock wave than in the untreated lymphedematous ears. Compared with the control group, shock wave treatment led to a significant decrease in the thickness of lymphedematous cars (3.80  $\pm$  0.25 mm vs 4.54  $\pm$  0.18 mm, P < .05). Immunohistochemistry for VEGFR3 showed the density of lymphatic vessels was significantly increased by shock wave

Research Paper

Journal of Vascular Research

J Vasc Res 2013;50:124-133 DOI: 10.1159/000343699

Accepted after revision: September 7, 2012 Published online: November 27, 2012

#### **Extracorporeal Shock Wave Therapy Combined with Vascular Endothelial Growth Factor-C Hydrogel for Lymphangiogenesis**

In Gul Kim<sup>a</sup> Ji Youl Lee<sup>a</sup> David S. Lee<sup>b</sup> Jeong Yi Kwon<sup>c</sup> Ji Hye Hwang<sup>c</sup>

<sup>a</sup>Department of Urology, Seoul St. Mary's Hospital, The Catholic University of Korea, <sup>b</sup>Yongsan International School, and Department of Physical and Rehabilitation Medicine, Sungkyunkwan University School of Medicine, Samsung Medical Center, Seoul, Korea

Eur J Vasc Endovasc Surg. 2011 Aug; 42(2):254-60. doi: 10.1016/j.ejvs.2011.02.029. Epub 2011 Mar 31.

#### Extracorporeal shock wave therapy induces therapeutic lymphangiogenesis in a rat model of secondary lymphoedema.

Serizawa F1, Ito K, Matsubara M, Sato A, Shimokawa H, Satomi S.

Author information

#### Abstract

OBJECTIVE: Lymphoedema is a common complication after cancer treatment. We have reported that low-energy extracorporeal shock wave (SW) therapy up-regulates vascular endothelial growth factor (VEGF) in ischaemic myocardium. As VEGF plays an important role in lymphangiogenesis, we investigated whether our low-energy SW therapy enhances lymphangiogenesis in rats.

METHODS: We created a tail model of lymphoedema in rats. The tail was treated with or without low-energy SW therapy (0.25 mJ mm(-2), 500 impulses) four times (days 3, 5, 7, and 9). The tail volume and the fluorescence intensity of indocyanine green (ICG) were measured. The expression of VEGF-C and basic fibroblast growth factor (bFGF) were evaluated by RT-PCR, and the lymphatic vessel density was assessed

RESULTS: The tail volume increased significantly in the control group and was significantly improved in the SW group. The lymphatic system function (evaluated with fluorescence intensity of ICG), the lymphatic vessel density, and the expression of VEGF-C and bFGF were all enhanced by the SW therapy (all P < 0.05).

CONCLUSIONS: The low-energy SW therapy induces therapeutic lymphangiogenesis by up-regulating VEGF-C and bFGF, and improves lymphoedema in a rat-tail model, suggesting that low-energy SW therapy could be a non-invasive and effective strategy for lymphoedema in

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PMID: 21454105 DOI: 10.1016/j.ejvs.2011.02.029

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# Klinische Studienlage



72 Patienten eingeschlossen (25 prim. LÖ, 47 sek. LÖ)

Alter: 70-85

### **Prozedere:**

10 Einheiten rESWT 2500-3500 Impulse (abh.von Größe der fibrotischen Areale)

Michelini et al. BMC Geriatrics 2010, 10(Suppl 1):A105 http://www.biomedcentral.com/1471-2318/10/S1/A105



#### MEETING ABSTRACT

**Open Acces** 

### Treatment of geriatrics lymphedema with shockwave therapy

S Michelini<sup>\*</sup>, M Cardone, A Failla, G Moneta, A Fiorentino, F Cappellino From de Senectute: Age and Health Forum Catanzaro, Italy. 5-7 December 2009



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### Evaluierung:

Baseline und 4 Wochen nach ESWT

### Ergebnis:

- Durchschn. Reduktion Extremitätenumfang von 26%
- Durchschn. Reduktion Hautdicke von 32% (hochauflöslicher US)
- Klinische Verbesserung der sklerosierten Hautareale



#### 7 Patientinnen

- mit fortgeschrittenen dermatofibrotischen Veränderungen
- kein Ansprechen auf KPE

#### **Prozedere:**

4x rESWT (2 Wochen), je 2000 Impulse

1000 Impulse "fibröseste" Läsion 1000 Impulse andere Läsionen

(2 Gruppen: 3 Pat. ESWT, 4 ESWT+KPE)

#### **Original Article**

Ann Rehabil Med 2013;37(2):229-234 pISSN: 2234-0645 • eISSN: 2234-0653 http://dx.dol.org/10.5535/arm.2013.37.2.229



#### Clinical Outcomes of Extracorporeal Shock Wave Therapy in Patients With Secondary Lymphedema: A Pilot Study

Hasuk Bae, MD, Ho Jeong Kim, MD

 $Department of Rehabilitation \ Medicine, Ewha \ Womans \ University \ School \ of \ Medicine, Seoul, Korean \ Grant \ Medicine, Seoul, Womans \ University \ School \ of \ Medicine, Seoul, Korean \ Grant \ Medicine, Seoul, Womans \ University \ School \ of \ Medicine, Seoul, Korean \ Grant \ Medicine, Seoul, Womans \ University \ School \ of \ Medicine, Womans \ University \ School \ of \ Medicine, Womans \ University \ School \ of \ Medicine, Womans \ University \ School \ of \ Medicine, Womans \ University \ School \ Of \ Medicine, Womans \ University \ Of \ Medicine, Womans \ University \ Of \ Medicine, Womans \ Of \ Medici$ 



### Evaluierung:

Baseline und unmittelbar nach Therapieende

- Umfangmessungen
- Volumetrie
- Hautdicke, Hauthärte, Sensible Defizite (VAS)

#### **Original Article**

Ann Rehabil Med 2013;37(2):229-234 pISSN: 2234-0645 • eISSN: 2234-0653 http://dx.doi.org/10.5535/arm.2013.37.2.229



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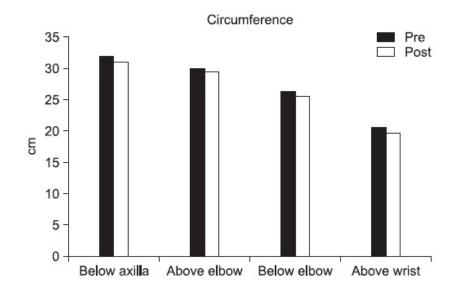


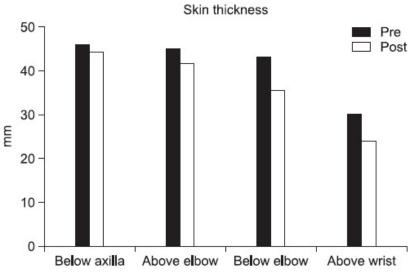




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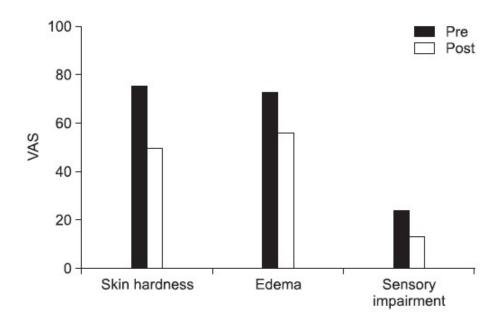






Clinical Outcomes of Extracorporeal Shock Wave Therapy in Patients With Secondary Lymphedema: A Pilot Study

nartment of Rehabilitation Medicine, Ewha Womans University School of Medicine, Seoul, Kor



**Gruppe ESWT+KPE:** 

+5% Volumenreduktion



11 PatientInnen mit sek.LÖ nach Brustkrebsbehandlung

#### **Procedere:**

12 (3x/wo.) rESWT-Einheiten

(2500 Impulse, 2 bar, 4 Hz)

Axilla 750 Impulse Cubita 250 Impulse Arm/Unterarm/Hand 1500 Impulse



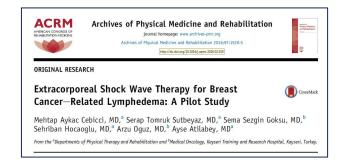
Characteristics	Value
Age (y)	50.63±7.03
Female	11
Lymphedema duration (mo)	12 (6-84)
Received chemotherapy	11
Received radiotherapy	8
Lymphedema severity at baseline	
Mild	1
Moderate	2
Severe	8
Lymphedema severity at 6mo after treatr	ment
Mild	3
Moderate	5
Severe	3



### Evaluierung:

baseline, posttherap., 1, 3 und 6 Mo.

- Volumetrische Vermessung
- QuickDASH





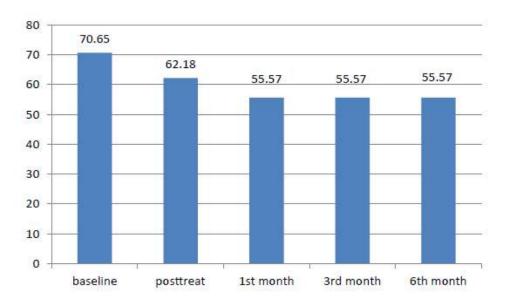


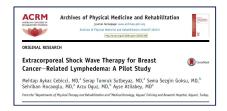
Fig 3 QuickDASH scores at baseline, posttreatment, and 1, 3, and 6 months after treatment.



#### volume (ml)



Fig 2 Mean volume displacement of the affected upper extremity at baseline, posttreatment, and 1, 3, and 6 months after treatment.





### Conclusio

- Stoßwellentherapie hat positive Effekte auf LÖ
- Technische Parameter sind zu klären
- Untersuchungen zum Wirkmechanismus notwendig
- Weiterführende Studien sind notwendig (höhere Fallzahl, kontrolliert, rand.)



### Kontakt:

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