



University of Groningen

20 year evolution of Glyaderm® dermal regeneration matrix

Pirayesh, Ali

DOI: 10.33612/diss.760333475

IMPORTANT NOTE: You are advised to consult the publisher's version (publisher's PDF) if you wish to cite from it. Please check the document version below.

Document Version Publisher's PDF, also known as Version of record

Publication date: 2023

Link to publication in University of Groningen/UMCG research database

Citation for published version (APA):

Pirayesh, A. (2023). 20 year evolution of Glyaderm® dermal regeneration matrix: the first non-commercial dermal regeneration matrix. [Thesis fully internal (DIV), University of Groningen]. University of Groningen. https://doi.org/10.33612/diss.760333475

Copyright

Other than for strictly personal use, it is not permitted to download or to forward/distribute the text or part of it without the consent of the author(s) and/or copyright holder(s), unless the work is under an open content license (like Creative Commons).

The publication may also be distributed here under the terms of Article 25fa of the Dutch Copyright Act, indicated by the "Taverne" license. More information can be found on the University of Groningen website: https://www.rug.nl/library/open-access/self-archiving-pure/taverneamendment.

If you believe that this document breaches copyright please contact us providing details, and we will remove access to the work immediately and investigate your claim.

Downloaded from the University of Groningen/UMCG research database (Pure): http://www.rug.nl/research/portal. For technical reasons the number of authors shown on this cover page is limited to 10 maximum.

CHAPTER 7

BURN SCAR ASSESSMENT -A SYSTEMATIC REVIEW OF OBJECTIVE SCAR ASSESSMENT TOOLS

BURNS (2010)

N. BRUSSELAERS A. PIRAYESH H. HOEKSEMA J. VERBELEN S. BLOT S. MONSTREY

Abstract

Purpose

All deep second and third degree burns are at risk to develop hypertrophic scars which can severely undermine the quality of survival. To assess the severity of scarring, several technical devices or tools have been introduced to evaluate one or more aspects of the scar, enabling comparison of different treatment protocols and allowing an objective follow-up. The objective of this study was to review which tools can be used in objective burn scar assessment.

Basic procedures

The Systematic literature search involving PubMed, the Web of Science (incl. Science Citation Index).

Main findings

51 articles with burn scar assessment as main topic were found. Several characteristics of the scar can be assessed, such as color, metric features and elasticity, but none of the available tools covers the whole aspect of the scar. Especially subjective factors such as pain and itching cannot be assessed with those tools, in spite of their great impact on the patient's quality of life.

Conclusions

Scar tools enable objective and reproducible evaluation of scars, which is essential for scientific studies and medico-legal purposes, and in selected cases for the clinical follow-up of an individual patient. Further studies to evaluate these tools on scars are nevertheless required.

Introduction

In the past decennia, major improvements in burn management have resulted in a substantially increased survival of severely burned patients¹⁻³. Unfortunately, this has not always been paralleled with a similar increase in quality of life for these patients^{4,5}. One of the major long-term problems in burn care is the formation of hypertrophic scars, which lead to aesthetical but also functional problems (e.g. contractures) and also cause a considerable psychological burden. Therefore, even the early burn treatment is guided and influenced by the risk of hypertrophic scar formation. The assessment of the natural healing potential is for example based on depth assessment, which also predicts the risk of abnormal scarring. The critical depth for excessive scar formation is in the deep dermis meaning that superficial burns should heal without leaving a scar, while deep burns always are 'at risk'⁶⁻⁹. Consequently, the longer the healing, the higher the risk of hypertrophic scarring¹⁰. The general rule in burn surgery is therefore to operate burns which will not heal within 2-3 weeks after the initial trauma^{7,11,12}.

To assess the severity of scarring, several scar scales have been developed over the last 30 years, of which the Vancouver Scar Scale and the POSAS scale (Patient & Observer Scar Assessment Scale) are the most widely used¹³⁻¹⁶. Scar scales include several variables such as color, extent and may even contain subjective factors such as pain and itching which are subject to major inter-patient variations. In general, scar scales are considered to be a subjective scoring system, because it is susceptible to important variation between different assessors (inter-assessor variation). To obtain a more objective evaluation of the scar, several devices or tools used in other medical specialties or even in the industry (e.g. assessment of textile color, elasticity of plastics) were introduced for the assessment of scars. These tools should provide a more objective and reliable evaluation of the scar, by a better reproducibility and lower inter-assessor variation.

In this overview we only focus on the scar tools, addressing the applied physical principles, and mentioning the most commonly described tools used for burn scar assessment.

Methods

Criteria for considering articles for inclusion

Articles dealing with non-invasive burn scar assessment with technical devices as a major topic were included. Scar scales without any technical analysis are excluded as well as histopathologic evaluations of scar biopsies. Articles comparing the influence of wound or scar treatments were also excluded.

Search methods

We conducted a systematic literature search involving PubMed and the Web of Science (which also contains major congress abstracts)^{17,18}. The Cochrane Library did

not contain relevant articles. We searched PubMed from 1960 until February 2009 (date of search 18 February), using the Medical Subject Headings (MeSH) 'Burns' and 'Cicatrix' (**Figure 1**). This search retrieved 1974 articles, whereof only articles with those terms as major topic were included, and the MeSH 'burns' were not exploded, excluding articles about chemical, electrical, eye and sun burns and inhalation injury (n = 928). Limits were set to English, French and Dutch articles about human studies (n = 597). The MeSH terms 'Surgical flaps' and 'Neoplasms' excluded irrelevant articles dealing with flap surgery (n = 78) and cancer development in burn scars (n = 91). From the remaining 428 articles, 67 articles were selected based on title and abstract, of which 30 articles corresponded with the inclusion and exclusion criteria¹⁹⁻⁴⁸.

We also searched the Web of Science on the terms 'scar', 'cicatrix', 'burns' or 'burn', but this search was too wide, because, even after language selection, it led to almost 44,000 hits. Therefore, more strict combinations were performed ('burn', 'burns', 'thermal injury', 'cicatrix' and 'scar'), leading to six additional articles^{15,49-53}.

After searching reference lists and Science Citation Index of the relevant articles, 15 additional articles were included⁵⁴⁻⁶⁸. We finally selected 51 articles, including six reviews and editorials.

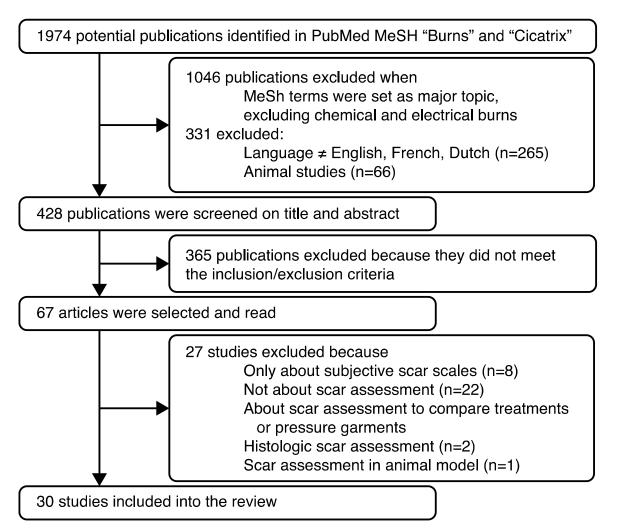


Figure 1. Flowchart of the systematic literature search (PubMed).

Quality assessment

Reproducibility of the assessments of these tools is evaluated (if described), and if possible the additional value of the device compared with visual assessment (e.g. with the scar scales) and/or other tools is discussed. The amount of articles (number of references) discussing a certain technique for scar assessment also reflects the relevance of implementing a similar device in clinical practice and its current popularity.

Results

Original articles, reviews and editorials dealing with burn scar assessment were retained. We chose to classify these tools into 4 groups based on the assessed variables: (A) color (vascularization, pigmentation), (B) metric variables (extent, height and volume), (C) biomechanical properties (e.g. elasticity, stiffness) and (D) physiologic changes (e.g. hydratation).

Colour evaluation

Color is probably the most complex characteristic of a scar and is mainly composed out of 3 components: the brown melanin pigment, the red oxyhemoglobin in the cutaneous vasculature (amount and oxygenation of blood vessels) and the yellow/ orange bile and carotene pigments³⁹. The thickness of the skin layers, the reflection from the skin surface (texture) and the circumstances (e.g. temperature, light) also influence the color perception^{38,39,69}. When using video or photographic images, these are strongly influenced by the settings of the camera (e.g. aperture, shutter-time) and the circumstances (e.g. light, temperature)⁵⁹.

Visual assessment is an effective but subjective way to evaluate color, with a considerable inter-observer variation¹⁵. Although the observer may distinguish thousands of colors, the human brain cannot reliably and accurately quantify the color or its intensity^{15,31,33,42}. Moreover, memorizing colors is difficult, complicating the quality of scar color ratings for follow-up³³. Therefore, several tools are developed to evaluate color in an objective and reproducible way, classified by the used principles: (1) reflection or absorption of light, (2) laser based methods and (3) computerized analysis of photographs.

Reflectance and absorption of light

It is not possible to obtain a perfect correlation between skin melanin or blood content and skin color³⁰. Therefore, the optimal method to assess skin color is not histologic or chemical but spectrophotmetric^{30,56,70,71}. Spectrometry is based on the reflectance and absorption of light and describes (i) the brightness and changes along the red-green and yellow-blue axis or (ii) the absorption of red and green light by melanin and hemoglobin respectively, resulting in the erythema and melanin index³⁰.

Different devices are used for color analysis of scars and skin diseases: (i) tristimulus colorimeters such as the Minolta Chromameter[®] (Konica), Labscan[®] (HunterLab) and the Micro Color (Dr. Lange GmbH), and the (ii) narrowband simple reflectance meters such as the DermaSpectrometer[®] (Cortex Technologies) and the Mexameter[®] (Courage&Khazaka)^{20,27,52,71}. These tools assess the vascularity and pigmentation better than scar scales and enable immediate 'on-site' evaluation²⁰.

Laser based methods

The laser based methods assess the bloodflow and apply red or near-infrared wavelengths³³. A considerably higher bloodflow is noted over immature burn scars, due to a higher vascularity. Structural changes may nevertheless interfere with perfusion measurements^{33,73}.

The Laser-Doppler Flowmeter (LDF) is used for the evaluation of cutaneous bloodflow to evaluate scar color^{15,30,42,59,60,66,72}. It measures the flow over a small location, limiting its value for extended, heterogeneous surfaces⁵⁹. This flowmeter is less sensitive than simple visual assessment of erythema and is therefore not recommended for color assessment⁶⁶. The Laser-Doppler Imaging (LDI) is a laserbeam which is used to scan several points across a tissue surface, generating a 2D color coded image directly related to the bloodflow^{19,31,33}. It is used for burn depth assessment^{7,74-77} but can also be used for scar evaluation, with fast and reproducible results^{19,31,33}. Another alternative is the Laser Speckle Imaging (LSPI) which uses digital image-processing techniques³¹. Moving red blood cells create dynamic interference patterns that change in time. The bloodflow maps are generated by coherent light reflected from stationary tissue producing a highly contrasted speckle pattern remaining static in time. LSPI allows for zooming in and increasing the resolution on a smaller field of view, in contrast to the LDI³¹.

Computer analysis of colour

Even standardized photographing fails to compare scars objectively when analyzed by the human brain^{30,66}. Therefore, several computer programs were developed to assess (digital) photographs^{59,60}. At first, color photographs were converted into black and white (BW), because of its less complex electronic make-up⁶⁰. The HSV-method analyzes three different aspects of color: the hue (dominant wave length e.g. red), the saturation (amount of white) and the value (amount of black), which are important in discriminating between colors, whereas in practice, differences in value mostly reflect varying levels of illumination^{66,78-82}. The colors can also be represented as combinations of the amount of red, green and blue ('primary colors') (RGB model)^{15,30,66}, or by the proportions of cyan, magenta and yellow ('secondary colors') and black (CMYK model). These 3 color models are equivalent and conversion between them is simple⁴¹. A card carrying standard colors (e.g. Pantone[®]) is used to frame the scar so that every picture would include areas of known color properties^{34,58}. Hereby, the influence of lightening conditions and camera settings can be subtracted, enabling an objective color evaluation^{30,38,41,65,66}.

Metric variables

Planimetry

Planimetry (or measuring surface area) is used to assess the extent of a scar and to detect contraction in time^{15,30,41,65,66}. The main problem is that scar margins become more difficult to delineate during scar maturation. Tracing these margins on clear plastic film and photography are most commonly applied⁶⁶. Photography is readily available, accurate and reliable (especially on flat or moderately curved surfaces), but standardized conditions are essential (distance, light, camera settings). Computer programs e.g. Image Tool[®] (C.C. Wilcox) can be used to determine the percentage of hypertrophic scars over the total scar area³⁰.

Height and volume assessment

Up to nine-fold increases in thickness have been described in scar tissue, but decreases can also be present⁴⁶. Hypertrophy and atrophy are quantified by measuring scar thickness or volume. The height of a scar can be evaluated subjectively but inaccurately, since the portion of the scar below the surface is not included⁴⁴. Some authors recommended histologic analysis of biopsied tissue (invasive technique), but skin biopsies may change in thickness when released from the tension and support provided in situ^{30,66}. It can also be questioned if the biopsy site is representative⁴⁴. Negative-positive moulds or replicas were used to make a 3D copy, accurately indicating height, extent and general appearance of a scar^{46,66,67,83}. This technique can be combined with photographs and tonometric assessment, and is also useful for evaluating the roughness^{46,60,66,84}. High frequency ultrasound (5-20MHz) tools such as the Dermascan[®] (Cortex Technology) provide reliable and accurate quantitative information on scar thickness^{22,26,35,40,44,48}. It is very sensitive in the localization of scar tissues, distinguishing them from normal skin, and for assessment of thickness and delineation of the extent of the scar^{40,44,65}. Therefore, portable devices e.g. TUPS (tissue ultrasound palpation system) were developed, facilitating clinical application. Although 3D ultrasound is available for clinical application, it is not widely used in scar evaluation because of its high costs²⁶. MRI has been used for the evaluation of normal skin but has not yet been applied on scars¹⁵.

Three dimensional techniques

Highly sophisticated, often expensive 3D methods became available for volume assessment, planimetry and analyzing roughness, including the use of full-body morphometric scanning, range scanners and 3D reconstructions (e.g. Vivid 900, Konica-Minolta and Vectra 3D imaging system, and Canfield Imaging Systems), but scientific studies assessing burn scars remain scarce^{30,41,54,64,65}. Advantages are the fast and direct, non-contact measurement of the surface and volume of the scar, macro- and micro-topometry, high resolution, high precision, and ease of handling^{60,85}. Range scanners project a light pattern onto a scene, which is

photographed by a regular camera³⁸. If the skin surface is uneven, which is the case in hypertrophic scarring, the projected light pattern appears distorted, which enables inferring the depths of points in a scene^{38,52}.

Biomechanical properties

Elasticity or stiffness

Several mechanisms can be used for evaluation of elasticity or stiffness. The elasticity of the skin is the property to return to its original shape when the stress is removed which caused deformation (e.g. external forces). Stiffness is the resistance of an elastic body to deformation by an applied force and can be quantified easier than elasticity. These methods described here originate from dermatology (e.g. Cutometer[®], Dermaflex[®] and Dermal torque meter[®]), ophthalmology (tonometers) and from industrial applications (durometer). These 'elastometers' can be classified by the applied biomechanical forces⁶⁶, which can be in a vertical direction: (i) suction or (ii) pressure; or horizontal: (iii) torsion or (iv) extension.

- 1. Suction methods: a controlled negative pressure is exerted over a small area of the scar, resulting in a skin deformation which is analyzed by a computer^{35,37,38,51,52,86}. The Cutometer[®] (Courage&Khazaka) proved to be highly reliable and reproducible for burn scars except for the most severe scars^{23,24,51,68,86}, but the size of the tool can be considered impractical³⁰. The Dermaflex[®] (Cortex Technology) is an alternative device with a larger diameter of the suction chamber (10 mm vs. 6 mm), but no scar assessment trials have been published yet^{49,66}.
- 2. Pressure methods or 'tonometers'^{38,66} originate from measuring intra-ocular pressure^{43,48} and hardness of metals and plastic^{57,87} and calculate the power required to produce a certain deformity⁴³. Several devices are developed and evaluated for skin elasticity measurement⁸⁷, of which several prototypes were tested on scars: cicatrometers⁴⁸, pneumatonometers^{30,62}, tonometers^{25,28,43} and durometers^{30,57}. They produce good results, but cannot be applied on scars above bone structures⁶⁶.
- 3. Torsion methods^{49,66} such as the Dermal Torque Meter[®] (Dia-Stron Ltd.) measure the torsion force needed to deform the skin. Only one scar study has been published, which reports resemblance with measurements with the Cutometer^{®49,66}.
- 4. Extension methods or 'extensioneters'⁶⁶ stretch the skin between two tabs to assess differences in extensibility or stiffness. This method has been described for scar evaluation^{38,50,66}, but scientific results are scarce⁶⁸.

Acoustic methods

Sound waves (5-8 kHz) are used to detect heterogeneity in the scar tissue, e.g. Shear Velocity Device, Reviscometer[®] (Courage&Khazaka)^{36,66}. A higher velocity (or speed) of wave transmission indicates a more dense structure (less deep penetration

of the waves), correlating with a higher degree of stiffness, related with scar contraction³⁶. These waves lie within the spectrum of normal hearing (which is 20Hz-20 kHz) and penetrate deeper in the skin than the ultrasound waves (5-20MHz), but both techniques have not been compared yet for scar evaluation³⁶.

Disability measuring

Because contractures primarily occur in joints, burn scars often compromise mobility. The mechanical impairment can be estimated by measuring the range of motion of a joint and is even included in some subjective scar assessment scales⁸⁸. The range of motion can be measured with goniometers (Greek for 'measuring an angle')^{66,89,90}. This term is used for simple plastic tools as well as computerized devices⁹¹.

It is also recommended to measure the disability itself (coordination, strength, skin sensibility)⁶⁶, e.g. by assessing daily life activities e.g. hand function⁶⁶. The faciometer[®] is an electronic device originally developed to assess the results after reconstructive surgery in cases of facial palsy. It consists of two calipers connected to a digital display, showing the actual distance between the calipers. Measurements of distances between specific stable and moving points are made at rest and after standardized maximal and submaximal (mimic) movements, enabling a 3D analysis⁵³. This tool proved to be useful for objective description of results after surgery for facial burns⁵³.

Pathophysiologic disturbances of the scar

Transcutaneous oxygen tension

Scar maturity has been related to transcutaneous oxygen tension^{15,52,55,61} which can nowadays be measured with electrodes on the skin (previously by subcutaneous needles)⁶¹. It is based on redox reactions occurring in the electrode modified by the inclusion of a heat source measuring the oxygen and the carbon dioxide that diffuses through the skin⁹². In hypertrophic scars the PO2 is lower than in healthy skin, but an increase is described which correlated with clinical improvement over 60 weeks of therapy^{55,61}. This technique seems to be abandoned for scar assessment, but is still used to assess limb ischemia^{92,93}.

Transepidermal water loss and moisture content

The skin acts as a barrier against permeation of external substances, as well as the water evaporation from the internal living tissue⁵². The water content in the skin preserves the softness and smoothness of the skin surface⁵², and this can be measured directly or by the transepidermal water loss (TEWL)^{52,56,94-100}. TEWL is strongly related with the moisture content of skin and can be measured with open or closed (insensitivity to external air currents) chamber systems. The open systems (e.g. Dermalab TEWL module, Tewameter[®], Courage&Khazaka) are the oldest and still

most widely used⁵². The advantages of the closed chamber systems (VapoMeter, Delphin Technologies) need to be determined⁹⁵.

Another method measures the hydratation of the skin surface (stratum corneum), which is directly proportional to retention of electrical charge, and can be measured by e.g. Skicon-2001 conductance meters (I.B.S. Co., Ltd), CorneoMeter[®] (Courage&Khazaka), and the DermaLab[®] (Cortex technology, also measures TEWL and elasticity)^{52,56,99,101,102}. These techniques are popular in the cosmetic industry, but also useful for evaluation of contact dermatitis and burn scars^{52,56,94}. Scar sites are dryer than control sites and seem to become dryer as they mature^{52,56,94,100}. The effect of environmental factors such as humidity (sweating) should be avoided, and showering and topical products are not allowed hours before measurement^{56,103}.

Discussion

Because hypertrophic scars are one of the major long-term problems after severe burns, scar prevention, treatment and assessment are of utmost importance. However, scar assessment is still a neglected area in the burn care, and a consensus about the ideal scar scale or tool is still lacking, probably due to the scarce amount of scientific studies. Several tools are currently promoted for (burn) scar evaluation, but these tools are mainly developed and commercialized for dermatologic use or for the cosmetic industry. Consequently, reference material for scar tissue is usually lacking, and no trials have been performed to compare the different tools for scar evaluation. Nevertheless, for most devices, the evaluation of skin or scar tissue should always be compared with a reference area of the patient, e.g. the other arm, because skin properties may vary considerably depending on the location on the body. The most important characteristics of the scar which can be analyzed by scar tools are the color, the thickness, the stiffness and the measurement of transepidermal water loss. These can all be assessed by different biomechanical techniques (sometimes combined in one device), with various degrees of complexity. The test results are preferable directly registered or integrated automatically in the computer system.

The question may rise what the therapeutic consequences are of evaluating scars, if you already use all preventive measures currently available. Nowadays, it is still difficult to predict which burn wound will certainly result in hypertrophic scarring and therefore preventive measures such as pressure therapy, splinting and silicones have become routine practice for all deep, extended burns in most burn units. Yet, it is useful to have an objective method to evaluate the degree of maturation of a scar, because it enables early adjustment of the therapy by introduction of extra preventive measures or earlier treatment e.g. by corticoid injections.

There is no doubt that objective scar assessment by scar tools definitely has an additional value in scientific studies, because different scars can be described and analyzed in more detail and compared mathematically. Therefore tools are statistically superior to scar scales and pictures; however the number of assessed variables is more limited than in scar scales. The role of scar tools in the daily clinical practice is less clear, because the tools are often large and expensive and increased

workload, time and costs. For this reason, scar scales are considered more costeffective and can also be used more easily in clinical practice (optimally combined with digital photographs). However, the scales are less objective than the tools due to the large inter- and intra-observer variation. In the (near) future, scar tools should become more accurate and reproducible than scales, and should detect derailment of the scar maturation earlier on, enabling earlier adjustment of therapy. Yet, at this stage, it is not possible to point out one ideal tool, and the optimal balance between accuracy, clinical and cost-economic applicability is still not reached within one single scar tool. Which tools will become more important will also be guided by the insight into the pathophysiology of scar formation. Recent studies report for example the major impact of the moisture content in scar maturation¹⁰⁴. Further comparative clinical trials are required to compare the reproducibility and accuracy of the scar tools.

To conclude, advances in technology resulted in several new promising techniques, but more scientific studies are needed before these scar tools can be implemented in the scientific and routine burn assessment. Besides scar tools, which can only assess a limited number of characteristics, an additional clinical evaluation will remain necessary, preferably by applying digital photography and a scar scale including the patient's perception of their scar (including pain, itching...) and the impact of the scar on the quality of life.

- 1. Development and validation of a model for prediction of mortality in patients with acute burn injury. Br J Surg 2009; 96: 111-7.
- 2. Brusselaers N, Juhasz I, Erdei I, Monstrey S, Blot S. Evaluation of mortality following severe burns injury in Hungary: external validation of a prediction model developed on Belgian burn data. Burns 2009;35:1009-14.
- 3. Brusselaers N, Hoste EA, Monstrey S, Colpaert KE, De Waele JJ, Vandewoude KH, et al. Outcome and changes over time in survival following severe burns from 1985 to 2004. Intensive Care Med 2005;31:1648-53.
- 4. Van Loey NE, Van Son MJ. Psychopathology and psychological problems in patients with burn scars: epidemiology and management. Am J Clin Dermatol 2003;4:245-72.
- 5. Brown BC, McKenna SP, Siddhi K, McGrouther DA, Bayat A. The hidden cost of skin scars: quality of life after skin scarring. J Plast Reconstr Aesthet Surg 2008;61:1049-58.
- 6. Hersch SJ. The early management of the burn wound and observations on hypertrophic scarring. With special reference to the deep dermal level and hypertrophic scarring. S Afr J Surg 1994;32:1-4.
- 7. Monstrey S, Hoeksema H, Verbelen J, Pirayesh A, Blondeel P. Assessment of burn depth and burn wound healing potential. Burns 2008;34:761-9.
- 8. Dunkin CS, Pleat JM, Gillespie PH, Tyler MP, Roberts AH, McGrouther DA. Scarring occurs at a critical depth of skin injury: precise measurement in a graduated dermal scratch in human volunteers. Plast Reconstr Surg 2007;119:1722-32. discussion 1733-4..
- 9. Bombaro KM, Engrav LH, Carrougher GJ, Wiechman SA, Faucher L, Costa BA, et al. What is the prevalence of hypertrophic scarring following burns? Burns 2003;29:299- 302.
- 10. Cubison TC, Pape SA, Parkhouse N. Evidence for the link between healing time and the development of hypertrophic scars (HTS) in paediatric burns due to scald injury. Burns 2006;32:992-9.
- 11. Brusselaers N, Lafaire C, Ortiz S, Jacquemin D, Monstrey S. The consensus of the surgical treatment of burn injuries in Belgium. Acta Chir Belg 2008;108:645-50.
- 12. Fraulin FO, Illmayer SJ, Tredget EE. Assessment of cosmetic and functional results of conservative versus surgical management of facial burns. J Burn Care Rehabil 1996;17:19-29.
- 13. Sullivan T, Smith J, Kermode J, McIver E, Courtemanche DJ. Rating the burn scar. J Burn Care Rehabil 1990;11:256-60.
- 14. Draaijers LJ, Tempelman FR, Botman YA, Tuinebreijer WE, Middelkoop E, Kreis RW, et al. The patient and observer scar assessment scale: a reliable and feasible tool for scar evaluation. Plast Reconstr Surg 2004;113:1960-5. discussion 1966-7.
- 15. Idriss N, Maibach HI. Scar assessment scales: a dermatologic overview. Skin Res Technol 2009;15:1-5.
- 16. Durani P, McGrouther DA, Ferguson MW. Current scales for assessing human scarring: a review. J Plast Reconstr Aesthet Surg 2009;62:713-20.
- 17. Moher D, Cook DJ, Eastwood S, Olkin I, Rennie D, Stroup DF. Improving the quality of reports of meta-analyses of randomised controlled trials: the QUOROM statement. Quality of reporting of meta-analyses. Lancet 1999;354:1896-900.
- Stroup DF, Berlin JA, Morton SC, Olkin I, Williamson GD, Rennie D, et al. Meta-analysis of observational studies in epidemiology: a proposal for reporting. Meta-analysis of Observational Studies in Epidemiology (MOOSE) group. JAMA 2000;283:2008-12.
- 19. Allely RR, Van-Buendia LB, Jeng JC, White P, Wu J, Niszczak J, et al. Laser Doppler imaging of cutaneous blood flow through transparent face masks: a necessary preamble to computer-controlled rapid prototyping fabrication with submillimeter precision. J Burn Care Res 2008;29:42-8.
- 20. Draaijers LJ, Tempelman FR, Botman YA, Kreis RW, Middelkoop E, van Zuijlen PP. Colour evaluation in scars: tristimulus colorimeter, narrow-band simple reflectance meter or subjective evaluation? Burns 2004;30:103-7.
- 21. Richard RL. Documenting changes in burn scars over time. J Burn Care Rehabil 2005;26:272.
- 22. Du YC, Lin CM, Chen YF, Chen CL, Chen T. Implementation of a burn scar assessment system by ultrasound techniques. Conf Proc IEEE Eng Med Biol Soc 2006;1:2328-31.
- 23. Nedelec B, Correa JA, Rachelska G, Armour A, LaSalle L. Quantitative measurement of hypertrophic scar: intrarater reliability, sensitivity, and specificity. J Burn Care Res 2008;29:489-500.
- 24. Nedelec B, Correa JA, Rachelska G, Armour A, LaSalle L. Quantitative measurement of hypertrophic scar: interrater reliability and concurrent validity. J Burn Care Res 2008;29:501-11.
- 25. Corica GF, Wigger NC, Edgar DW, Wood FM, Carroll S. Objective measurement of scarring by multiple assessors: is the tissue tonometer a reliable option? J Burn Care Res 2006;27:520-3.
- 26. Lau JC, Li-Tsang CW, Zheng YP. Application of tissue ultrasound palpation system (TUPS) in objective scar evaluation. Burns 2005;31:445-52. 8.
- 27. Li-Tsang CW, Lau JC, Liu SK. Validation of an objective scar pigmentation measurement by using a spectrocolorimeter. Burns 2003;29:779-84. 42.
- 28. Lye I, Edgar DW, Wood FM, Carroll S. Tissue tonometry is a simple, objective measure for pliability of burn scar: is it reliable? J Burn Care Res 2006;27:82-5.
- 29. Masters M, McMahon M, Svens B. Reliability testing of a new scar assessment tool, Matching Assessment of Scars and Photographs (MAPS). J Burn Care Rehabil 2005;26:273-84.
- 30. Oliveira GV, Chinkes D, Mitchell C, Oliveras G, Hawkins HK, Herndon DN. Objective assessment of burn scar vascularity, erythema, pliability, thickness, and planimetry. Dermatol Surg 2005;31:48-58.

- 31. Stewart CJ, Frank R, Forrester KR, Tulip J, Lindsay R, Bray RC. A comparison of two laser-based methods for determination of burn scar perfusion: laser Doppler versus laser speckle imaging. Burns 2005;31:744-52.
- 32. Zhang Y, Goldgof DB, Sarkar S, Tsap LV. A modeling approach for burn scar assessment using natural features and elastic property. IEEE Trans Med Imaging 2004;23:1325-9.
- 33. Bray R, Forrester K, Leonard C, McArthur R, Tulip J, Lindsay R. Laser Doppler imaging of burn scars: a comparison of wavelength and scanning methods. Burns 2003;29:199- 206.
- 34. Davey RB, Sprod RT, Neild TO. Computerised colour: a technique for the assessment of burn scar hypertrophy. A preliminary report. Burns 1999;25:207-13.
- 35. Fong SS, Hung LK, Cheng JC. The cutometer and ultrasonography in the assessment of postburn hypertrophic scar—a preliminary study. Burns 1997;23(Suppl. 1):S12-8.
- McHugh AA, Fowlkes BJ, Maevsky EI, Smith Jr DJ, Rodriguez JL, Garner WL. Biomechanical alterations in normal skin and hypertrophic scar after thermal injury. J Burn Care Rehabil 1997;18:104-8.
- 37. Nedelec B, Shankowsky HA, Tredget EE. Rating the resolving hypertrophic scar: comparison of the Vancouver Scar Scale and scar volume. J Burn Care Rehabil 2000;21:205-12.
- 38. Powers PS, Sarkar S, Goldgof DB, Cruse CW, Tsap LV. Scar assessment: current problems and future solutions. J Burn Care Rehabil 1999;20:54-60. discussion 53.
- 39. Tyack ZF, Pegg S, Ziviani J. Postburn dyspigmentation: its assessment, management, and relationship to scarring—a review of the literature. J Burn Care Rehabil 1997;18:435-40.
- 40. Van den Kerckhove E, Staes F, Flour M, Stappaerts K, Boeckx W. Reproducibility of repeated measurements on post-burn scars with Dermascan C. Skin Res Technol 2003;9:81-4.
- 41. Wood FM, Currie K, Backman B, Cena B. Current difficulties and the possible future directions in scar assessment. Burns 1996;22:455-8.
- 42. Ehrlich HP, Kelly SF. Hypertrophic scar: an interuption in the remodeling of repair—a Laser Doppler blood flow study. Plast Reconstr Surg 1992;90:993-8.
- 43. Esposito G, Ziccardi P, Scioli M, Pappone N, Scuderi N. The use of a modified tonometer in burn scar therapy. J Burn Care Rehabil 1990;11:86-90.
- 44. Hambleton J, Shakespeare PG, Pratt BJ. The progress of hypertrophic scars monitored by ultrasound measurements of thickness. Burns 1992;18:301-7.
- 45. Leung KS, Sher A, Clark JA, Cheng JC, Leung PC. Microcirculation in hypertrophic scars after burn injury. J Burn Care Rehabil 1989;10:436-44.
- 46. Sawada Y. A method of recording and objective assessment of hypertrophic burn scars. Burns 1994;20:76-8
- 47. Chu BM, Brody G. Nondestructive measurements of the properties of healing burn scars. Med Instrum 1975;9:139-42
- 48. Katz SM, Frank DH, Leopold GR, Wachtel TL. Objective measurement of hypertrophic burn scar: a preliminary study of tonometry and ultrasonography. Ann Plast Surg 1985;14:121-7.
- 49. Boyce ST, Supp AP, Wickett RR, Hoath SB, Warden GD. Assessment with the dermal torque meter of skin pliability after treatment of burns with cultured skin substitutes. J Burn Care Rehabil 2000;21:55-63.
- 50. Clark JA, Cheng JC, Leung KS. Mechanical properties of normal skin and hypertrophic scars. Burns 1996;22:443-6.
- 51. Dobrev HP. A study of human skin mechanical properties by means of Cutometer. Folia Med (Plovdiv) 2002;44:5-10.
- 52. Kim YJ, Kim MY, Lee PK, Kim HO, Park YM. Evaluation of natural change of skin function in split-thickness skin grafts by noninvasive bioengineering methods. Dermatol Surg 2006;32:1358-63.
- 53. Koller R, Kargul G, Giovanoli P, Meissl G, Frey M. Quantification of functional results after facial burns by the faciometer. Burns 2000;26:716-23.
- 54. Ardehali B, Nouraei SA, Van Dam H, Dex E, Wood S, Nduka C. Objective assessment of keloid scars with threedimensional imaging: quantifying response to intralesional steroid therapy. Plast Reconstr Surg 2007;119:556-61.
- 55. Berry RB, Tan OT, Cooke ED, Gaylarde PM, Bowcock SA, Lamberty BG, et al. Transcutaneous oxygen tension as an index of maturity in hypertrophic scars treated by compression. Br J Plast Surg 1985;38:163-73.
- 56. Ho DQ, Bello YM, Grove GL, Manzoor J, Lopez AP, Zerweck CR, et al. A pilot study of noninvasive methods to assess healed acute and chronic wounds. Dermatol Surg 2000;26:42-9.
- 57. Magliaro A, Romanelli M. Skin hardness measurement in hypertrophic scars. Wounds 2003;15:66-70.
- 58. Powell MW, Sarkar S, Goldgof DB, Ivanov K. A methodology for extracting objective color from images. IEEE Trans Syst Man Cybern B Cybern 2004;34:1964-78.
- 59. Roques C, Curtet A, Druilhe E, Prieur F, Redlinger A, Guoyt L, et al. E' valuation de la couleur des cicatrices de brûlures par vision artificielle: étude préliminaire. Brûlures 2002;3:16-20.
- 60. Roques C, Teot L. A critical analysis of measurements used to assess and manage scars. Int J Low Extrem Wounds 2007;6:249-53.
- 61. Sloan DF, Brown RD, Wells CH, Hilton JG. Tissue gases in human hypertrophic burn scars. Plast Reconstr Surg 1978;61:431-6.
- 62. Spann K, Mileski WJ, Atiles L, Purdue G, Hunt J. The 1996 Clinical Research award. Use of a pneumatonometer in burn scar assessment. J Burn Care Rehabil 1996;17:515-7. Burns 36 (2010) 1157-1164 1163

- 63. Suetake T, Sasai S, Zhen YX, Ohi T, Tagami H. Functional analyses of the stratum corneum in scars. Sequential studies after injury and comparison among keloids, hypertrophic scars, and atrophic scars. Arch Dermatol 1996;132:1453-8.
- 64. Taylor B, McGrouther DA, Bayat A. Use of non-contact 3D digitaliser to measure the volume of keloid scars: a useful tool for scar assessment? J Plast Reconstr Aesthet Surg 2007;60:87-94.
- 65. Tsap LV, Goldgof DB, Sarkar S, Powers PS. A vision-based technique for objective assessment of burn scars. IEEE Trans Med Imaging 1998;17:620-33.
- 66. van Zuijlen PP, Angeles AP, Kreis RW, Bos KE, Middelkoop E. Scar assessment tools: implications for current research. Plast Reconstr Surg 2002;109:1108-22.
- 67. Wang ZY, Zhang J, Lu SL. Objective evaluation of burn and post-surgical scars and the accuracy of subjective scar type judgment. Chin Med J (Engl) 2008;121:2517-20.
- 68. Bartell TH, Monafo WW, Mustoe TA. A new instrument for serial measurements of elasticity in hypertrophic scars. J Burn Care Rehabil 1988;9:957-60.
- 69. Argenbright LW, Forbes PD. Erythema and skin blood content. Br J Dermatol 1975;106:569-74.
- 70. van Zuijlen PP, Vloemans JF, van Trier AJ, Suijker MH, van Unen E, Groenevelt F, et al. Dermal substitution in acute burns and reconstructive surgery: a subjective and objective long-term follow-up. Plast Reconstr Surg 2001;108:1938-46.
- 71. Clarys P, Alewaeters K, Lambrecht R, Barel AO. Skin color measurements: comparison between three instruments: the Chromameter(R), the DermaSpectrometer(R) and the Mexameter(R). Skin Res Technol 2000;6:230-8.
- 72. Leung KS, Cheng JC, Leung YK, Clark JA, Ma GF, Leung PC. In vivo study of the mechanical property of postburn hypertrophic scar tissues. J Burn Care Rehabil 1984;5:458-62.
- 73. Page RE, Robertson GA, Pettigrew NM. Microcirculation in hypertrophic burn scars. Burns Incl Therm Inj 1983;10:64-70.
- 74. Hoeksema H, Van de Sijpe K, Tondu T, Hamdi M, Van Landuyt K, Blondeel P, et al. Accuracy of early burn depth assessment by laser Doppler imaging on different days post burn. Burns 2009;35:36-45.
- 75. Pape SA, Skouras CA, Byrne PO. An audit of the use of laser Doppler imaging (LDI) in the assessment of burns of intermediate depth. Burns 2001;27:233-9.
- 76. Droog EJ, Steenbergen W, Sjoberg F. Measurement of depth of burns by laser Doppler perfusion imaging. Burns 2001;27:561-8.
- 77. Niazi ZB, Essex TJ, Papini R, Scott D, McLean NR, Black MJ. New laser Doppler scanner, a valuable adjunct in burn depth assessment. Burns 1993;19:485-9.
- 78. Bohannon RW, Pfaller BA. Documentation of wound surface area from tracings of wound perimeters. Clinical report on three techniques. Phys Ther 1983;63:1622-4.
- 79. Fuller FW, Mansour EH, Engler PE, Shuster B. The use of planimetry for calculating the surface area of a burn wound. J Burn Care Rehabil 1985;6:47-9.
- 80. Johnson M, Miller R. Measuring healing in leg ulcers: practice considerations. Appl Nurs Res 1996;9:204-8.
- 81. Kantor J, Margolis DJ. Efficacy and prognostic value of simple wound measurements. Arch Dermatol 1998;134:1571-4.
- 82. Langemo DK, Melland H, Hanson D, Olson B, Hunter S, Henly SJ. Two-dimensional wound measurement: comparison of 4 techniques. Adv Wound Care 1998;11:337-43.
- 83. Nedelec B, Ghahary A, Scott PG, Tredget EE. Control of wound contraction. Basic and clinical features. Hand Clin 2000;16:289-302.
- 84. Barbenel JC, Makki S, Agache P. The variability of skin surface contours. Ann Biomed Eng 1980;8:175-82.
- 85. Roques C, Teot L, Frasson N, Meaume S. PRIMOS: an optical system that produces three-dimensional measurements of skin surfaces. J Wound Care 2003;12:362-4.
- 86. Draaijers LJ, Botman YA, Tempelman FR, Kreis RW, Middelkoop E, van Zuijlen PP. Skin elasticity meter or subjective evaluation in scars: a reliability assessment. Burns 2004;30:109-14.
- 87. Falanga V, Bucalo B. Use of a durometer to assess skin hardness. J Am Acad Dermatol 1993;29:47-51.
- 88. Smith AC, Kimble R, Mill J, Bailey D, O'Rourke P, Wootton R. Diagnostic accuracy of and patient satisfaction with telemedicine for the follow-up of paediatric burns patients. J Telemed Telecare 2004;10:193-8.
- 89. Low JL. The reliability of joint measurement. Physiotherapy 1976;62:227-9.
- 90. Boone DC, Azen SP, Lin CM, Spence C, Baron C, Lee L. Reliability of goniometric measurements. Phys Ther 1978;58:1355-90.
- 91. Harvey KD, Barillo DJ, Hobbs CL, Mozingo DW, Fitzpatrick JC, Cioffi WG, et al. Computer-assisted evaluation of hand and arm function after thermal injury. J Burn Care Rehabil 1996;17:176-80. discussion 175.
- 92. Rodrigues LM, Roberto MA. Characterization strategies for the functional assessment of the cutaneous lesion. Burns 2006;32:797-801.
- 93. Carter SA, Tate RB. The relationship of the transcutaneous oxygen tension, pulse waves and systolic pressures to the risk for limb amputation in patients with peripheral arterial disease and skin ulcers or gangrene. Int Angiol 2006;25:67-72.
- 94. Shah JH, Zhai H, Maibach HI. Comparative evaporimetry in man. Skin Res Technol 2005;11:205-8.

- 95. Cohen JC, Hartman DG, Garofalo MJ, Basehoar A, Raynor B, Ashbrenner E, et al. Comparison of closed chamber and open chamber evaporimetry. Skin Res Technol 2009;15:51-4.
- 96. Nuutinen J, Alanen E, Autio P, Lahtinen MR, Harvima I, Lahtinen T. A closed unventilated chamber for the measurement of transepidermal water loss. Skin Res Technol 2003;9:85-9.
- 97. Gioia F, Celleno L. The dynamics of transepidermal water loss (TEWL) from hydrated skin. Skin Res Technol 2002;8:178-86.
- 98. André, De Wan M, Lefèvre P, Thonnard J-L. Moisture evaluator: a direct measure of fingertip skin hydratation during object manipulation. Skin Res Technol 2008;14:385-9.
- 99. Alanen E, Nuutinen J, Nicklén K, Lahtinen T, Mönkkönen J. Measurement of hydratation in the stratum corneum with the MoistureMeter and comparison with the Corneometer. Skin Res Technol 2004;10:32-7.
- 100. Suetake T, Sasai S, Zhen YX, Ohi T, Tagami H. Functional analyses of the stratum corneum in scars. Arch Dermatol 1996;132:1453-8.
- 101. Agache P, Mary S, Muret P, Matta AM, Humbert P. Assessment of the water content of the stratum corneum using a sorption-desorption test. Dermatology 2001;202:308-13.
- 102. Tagami H, Masatoshi O, Keije I, Kanamura Y, Yamada M, Ichijo B. Evaluation of the skin surface hydratation in vivo by electrical measurement. J Invest Dermatol 1980;75:500-7.
- 103. Tagami H. In: Elsner P, Berardesca E, Maibach H, editors. Hardware and measuring principle: skin conductance. Bioengineering of skin: water and the stratum corneum. Boca Raton, FL: CRC Press; 1994. p. 197-203.
- 104. Mustoe TA. Evolution of silicone therapy and mechanism of action in scar management. Aesthetic Plast Surg 2008;32:82-92.