Vascular complications Part 1: Prevention

BY JANI VAN LOGHEM, PIETER SIEBENGA AND JOB THUIS

Vascular complications from the use of soft tissue fillers can be devastating. In the first of a two-part series, the authors discuss how to avoid these adverse events.

n the past decade, the number of aesthetic procedures with soft tissue fillers (STF) has increased dramatically. Although the outcome profile is mostly favourable, the number of adverse events (AEs) is increasing as STF use becomes more commonplace. The vast majority of STF-associated AEs are transient and mild in intensity and need little medical intervention. However, vascular complications are considered as one of the most severe, yet rare, complications following treatment with STF and require immediate and appropriate medical treatment. Depending on the extent of the intravascular embolus, the clinical effect can range from asymptomatic to devastating and life changing. Blindness is considered one of the most severe AEs after STF injections and can occur after a STF enters one of the arterial connections between the central retinal artery and the extraorbital arteries (Table 1). While the incidence of vascular compromise remains low, more cases are appearing, mainly due to the increased use of STF and more advanced indications being performed by a greater number of practitioners worldwide [1-3]. Two main types of ischaemia are discussed in this article: retinal ischaemia (potentially leading to blindness) and peripheral ischaemia (potentially leading to tissue necrosis and scarring).

Vascular compromise

Vascular compromise is the disruption of blood supply to an artery or vein after STF treatment. It can happen in any area of the face (or body) and leads - in the case of peripheral ischaemia - to a sudden onset of blanching, livedo reticularis and / or blue-grayish discoloration. Accompanying symptoms may include pain, aphasia or even hemiparesis. Vascular compromise of the retinal artery leads to a sudden onset of simultaneous occurrence of vision loss and eye pain. Ophthalmoparesis, ptosis, vertigo and fainting may occur simultaneously, as well as the above-mentioned peripheral ischaemic symptoms [4,5]. It is critical to recognise these as soon as possible so that treatment of the complication can be started immediately; there is only a short window of opportunity, often quoted as 60 minutes, before blindness becomes irreversible [6] with some recent reports of a shorter period.

Direct arterial embolisation may occur in an anterograde or retrograde fashion. In anterograde vascular occlusion, the injection pressure does not exceed arterial pressure, whereas the opposite is true for retrograde vascular occlusion. Anterograde vascular occlusion causes a decreased blood flow downstream to the vascular branches, resulting mainly in surface changes of the skin [7]. Retrograde vascular occlusion allows the STF to travel against the arterial blood flow. The product often travels back to a vascular bifurcation, at which point the arterial pressure gradient stabilises, and the embolus can then subsequently cause anterograde occlusion at a more distal location than the initial injection site [7]. The arterial occlusion, and subsequent

non-perfusion, is most likely to occur within distal capillaries, as demonstrated in a mouse animal model [8]. Apart from arterial embolisation, venous obstruction is a possible cause of vascular occlusion. However, as veins increase in diameter downstream of the capillary bed, arterial occlusion seems more relevant. Arterial compression is a theoretical possibility for obstruction of blood flow, but as evidence for this possible pathogenetic pathway is absent, the current article will focus on arterial embolisation.

Anatomy

A thorough understanding of peripheral arterial anatomy is essential to reduce the risk of both retinal and peripheral ischaemia. General statements can be made concerning the expected anatomical layer and the general direction of the arteries. Detailed anatomy of the facial vasculature is described elsewhere [9].

In a large global review of blindness secondary to facial filler injections, Beleznay et al. reported a total of 146 cases, although this inevitably represents under-reporting [10,11]. The most common locations for filler injections causing vision changes were the nasal region (n=52), the glabella (n=51), the forehead (n=21) and the nasolabial fold (NLF) (n=20) [10,11]. Less common sites were the temple, cheek, chin and upper eyelid [11]. Historically, autologous fat injections caused the most visual compromise complications, attributed to the larger volumes and needle

Table 1: The facial branches of the ophthalmic artery and their main connections. Please note that these arteries also anastomose. Anatomical variations may exist.

Facial branch of the ophthalmic artery	Connecting to arteries in the face
Supratrochlear artery	Central forehead artery
Supraorbital artery	Superficial temporal artery
Dorsal nasal artery	Angular artery Lateral nasal artery Columellar artery
Lacrimal artery	Deep temporal artery (via zygomatical temporal artery) Zygomaticofacial artery
External nasal artery	Nasal arteries

bore size necessary. However, for the years 2015 to 2018, 81.3% of visual impairment / loss caused by SFT occurred with the injection of a hyaluronic acid. This probably reflects the increase in popularity of hyaluronic acid during this period [11].

Vascular compromise to the retina can result in permanent blindness if the retinal artery is occluded. Due to the short distance to the central retinal artery, injections of SFT in the glabellar region are most likely to cause vascular occlusion. In particular, the supratrochlear artery, which is itself a terminal branch of the ophthalmic artery, can be inadvertently accessed due to its superomedial location at the orbit and its extensive vascular anastomoses throughout the forehead and with the adjacent supraorbital artery [7].

Embolisation of the posterior ciliary arteries, which provide a blood supply to the optic nerve, may cause optic neuropathy. Lastly, occlusion of the middle cerebral artery, a branch of the internal carotid artery, can occur with significant injection pressure and result in cerebral infarction [11].

Injection techniques can be adjusted to reduce the risk of vascular complications (Table 2). Large boluses and the use of needles should be avoided [4,5]. Positioning the bevel of the needle or cannula in a different anatomical layer to that where the artery is most likely to be found, and injecting perpendicular to the expected direction of an artery, can also reduce the risk of injecting large amounts intravascularly.

Prevention

Although the risks of STF-associated vascular compromise cannot be eliminated completely, there are multiple steps a physician can take to avoid such complications. Firstly, the patient must be well informed about the risks of the treatment, and informed consent must be obtained before any procedure begins. Physicians are also advised to assemble an 'emergency tool kit' for the management of complications should they arise [12]. During the treatment, there are several recommendations to further reduce the risk of vascular filler complications; the 10 most important are listed below:

 Use blunt cannulas with a gauge of 25G or thicker (if the injector is comfortable with cannulas).

- Choose the anatomical layer with the lowest risk of arterial presence (Table 2).
- Choose a cannula direction as perpendicular to the artery as possible (Table 2).
- To pass a fascia, ligament or other expected resistance, some force is allowed.
- After reaching the desired anatomical layer, no force should be used ('tea-time technique') [13].
- When the cannula tip reaches a resistance, a gentle, back and forth motion, combined with a rolling technique (the 'pencil-rolling technique') should be used to pass resistances without using force [13].
- Take extra care near neuromuscular bundles, scars and other areas where arteries are surrounded with dense tissue that may prevent the artery from being pushed aside.
- Around the supratrochlear and dorsal nasal arteries, use the non-dominant thumb and index finger to press on the supratrochlear arteries to temporarily block blood flow (and prevent embolus displacement).

Table 2: The most important arteries in the face with their expected anatomical layer (based on Mendelson's 5-layer classification: 1: skin; 2: subcutaneous; 3: musculo-aponeurotic; 4: retaining ligaments and spaces; 5: periosteum and deep fascias) and their expected general direction. Note: anatomic variation is very common in arterial anatomy. IFS: inferior frontal septum; MFS: middle frontal septum; UFS: upper frontal septum.

Artery	Expected anatomical layer	Expected direction
Supratrochlear a.	Below the IFS: Level 2-5 Between IFS and MFS: Level 2-3 Above UFS: Level 2	Vertical
Supraorbital a.	Below the IFS: Level 2-5 Between IFS and MFS: Level 2-3 Above UFS: Level 2	Vertical with the lateral branch diagonal
Superficial temporal a.	Level 3	Parietal branch: vertical frontal branch: diagonal
Deep temporal a. (anterior and posterior)	Level 5	Vertical
Dorsal nasal a.	Level 2	Vertical
Lateral nasal a.	Level 2	Around the ala (vertical to horizontal)
Columellar a.	Level 2	Vertical
Angular a.	Level 2	Vertical
Facial a.	Below zygomaticus mm: level 4 Above zygomaticus mm: level 2	Below mandible: horizontal Between mandible and ala: diagonal
External carotid a.	Level 4	Vertical
Transverse facial a.	Level 4	Horizontal
Superior labial a.	Level 4	Horizontal
Inferior labial a.	Level 4	Lateral: horizontal Medial: vertical
Submental artery	Level 4	Below mandible: horizontal At mentum: vertical
Infraorbital a.	Level 4-5	Diagonal, horizontal and vertical branches
Zygomaticofacial a.	Level 4-5	Diagonal, horizontal and vertical branches
Mental a.	Level 4-5	Diagonal, horizontal and vertical branches

- Inject low volumes (0.025ml or less per retrograde linear thread) to reduce the chance of reaching the central retinal artery (the JVL Bolus).
- Inject slowly to reduce chances of retrograde displacement of the embolus against the blood flow of the ophthalmic artery branches.

Aspiration is often listed as a prevention measure, but is influenced by needle diameter, needle length and the rheological properties of the STFs. One study has reported a true positive result in only 33% of the tests performed with STFs within one second of aspiration [14].

The JVL Bolus

In workshops focusing on rhinomodulation, Dr Fernando Silikovic (Buenos Aires, Argentina) incorporated the advice of the authors and used a maximum of 0.025ml per bolus or retrograde linear thread, and has named this small aliquot of filler 'The JVL Bolus' [personal communication between J van Loghem and F Silikovic]. Cadaver studies show that only a small amount of product is required to travel from the external part of the supratrochlear artery inside the orbit to the ophthalmic artery, and thereby enter the central retinal artery at its bifurcation with the ophthalmic artery. The average volume measured was 0.085ml with a spread of 0.04-0.12ml [7]. This study was limited by the fact that data were obtained from only six fresh tissue cadaver heads (12 eyes). In addition, it cannot be assumed that findings from cadaveric head vasculature will perfectly simulate conditions of in vivo active human circulation (i.e. flow characteristics, temperature), which may account for some degree of volume discrepancy [7]. However, in light of these findings, extra safety precautions are desirable. Most commercially available syringes have markings on them, with the smallest being 0.05ml. As a practical solution, the authors advise using one marking for two boluses or retrograde linear threads, so that an amount of approximately 0.025ml is injected per bolus or retrograde linear thread (the JVL Bolus).

Using US as a preventative measure to avoid intravascular injection of soft tissue fillers

Recently, reports have been published promoting the use of doppler ultrasound (US) scans [15] prior to injection of STF. The rationale is that due to high arterial anatomic variations, the exact location of arteries remains unknown until they can actually be visualised. This technique may allow large diameter arteries to be localised. However, the only appropriate way for US to be used as a preventative tool would be to inject under US visualisation, as a difference of 1mm could mean the difference between intra- and extra-arterial injection.

There are some disadvantages associated with the use of US as a preventative tool for the avoidance of intravascular injection:

- The injector may have a false sense of security: the smaller diameter arteries cannot be seen due to the limited resolution of most devices. As we have learnt from cadaver dissections with dye-coloured arteries, the facial tissues are supplied by thousands of smaller diameter arteries that interconnect with the arterial system. In his article on management of vascular complications, Delorenzi compared the arterial network to an interstate highway system [16]. Even the smallest dirt roads eventually connect to the interstate highway and the driver can reach any destination. Similarly, in the face, injecting in a smaller diameter artery, invisible on the US screen, can lead to widespread necrosis, just as injecting in larger arteries can.
- Smaller needles cannot be used as injection under visualisation is virtually impossible with thin and short needles. The thinner needles (27-30G, 13mm) that are generally used for HA injection are difficult to see under US. Even more importantly, these needles are usually introduced perpendicular to the skin, which is not an ideal angle for US visualisation. If the needles enter at a smaller angle to the skin, their short length makes deep injections impossible.
- The non-dominant hand is unavailable for guiding the injection. Many expert injectors agree that the non-dominant hand is at least as important as the injecting hand. Often, the non-dominant hand is essential to advance the cannula or needle to a certain target area. Therefore, the use of US-guided injections may reduce the aesthetic outcome of soft tissue treatments.
- There are sterility issues: the injector has to use sterile US gel and a sterile protective rubber condom around the probe, as well as ensure that the gel itself does not enter the skin through the needle holes.

The main advantage of US as a preventative measure is that the injector can form a general idea of the location of the arteries. It is important that injectors using US preventatively are aware of the limited sensitivity of this safety test and practise the suggested technical preventative measures mentioned above.

References

- Hanke CW, Redbord KP. Safety and efficacy of poly-Llactic acid in HIV lipoatrophy and lipoatrophy of aging. J Drugs Dermatol 2007;6(2):123-8.
- Kadouch JA. Calcium hydroxylapatite: a review on safety and complications. J Cosmet Dermatol 2017;16(2):152-61.
- 3. Rayess HM, Svider PF, Hanba C, et al. A cross-sectional analysis of adverse events and litigation for injectable fillers. *JAMA Facial Plast Surg* 2018;**20(3)**:207-14.
- Van Loghem JAJ, Funt D, Pavicic T, et al. Managing intravascular complications following treatment with calcium hydroxylapatite: An expert consensus. J Cosmet Dermatol 2020;00:1-14.
- DeLorenzi C. Complications of injectable fillers, part 2: vascular complications. *Aesthet Surg J* 2014;34:584-600.
- Loh KT, Chua JJ, Lee HM, et al. Prevention and management of vision loss relating to facial filler injections. *Singapore Med J* 2016; 57:438-43.
- Khan TT, Colon-Acevedo B, Mettu P, et al. An anatomical analysis of the supratrochlear artery: considerations in facial filler injections and preventing vision loss. *Aesthet Surg J* 2017;**37(2)**:203-8.
- Chang S, Yousefi S, Qin J, et al. External compression versus intravascular injection: A mechanistic animal model of filler-induced tissue ischemia. *Ophthal Plast Reconstr Surg* 2016;32(4):261-6.
- Von Arx T, Tamura K, Yukiya O, et al. The Face A Vascular Perspective. A literature review. Swiss Dent J 2018;128(5):382-92.
- Beleznay K, Carruthers JD, Humphrey S, et al. Avoiding and Treating Blindness From Fillers: A Recent Review of the World Literature. *Dermatol Surg* 2015;41(10):1097-117.
- Beleznay K, Carruthers JD, Humphrey S, et al. Update on Avoiding and Treating Blindness From Fillers: A Recent Review of the World Literature. *Aesthet Surg J* 2019;**39(6)**:662-74.
- Signorini M, Liew S, Sundaram H, et al. Global Aesthetics Consensus: Avoidance and Management of Complications from Hyaluronic Acid Fillers— Evidence- and Opinion-Based Review and Consensus Recommendations. *Plast Reconstr Surg* 2016;**137**:961e.
- Van Loghem J. The Linear Bolus Technique and other practical tips for the advanced use of non-traumatic cannulas in aesthetic medicine. *PMFA News* 2016;**3(2)**: 12-3.
- Van Loghem JAJ, Fouché JJ, Thuis J. Sensitivity of aspiration as a safety test before injection of soft tissue fillers. J Cosmet Dermatol 2018;17(1):39-46.
- Schelke LW, Velthuis P, Kadouch J, Swift A. Early ultrasound for diagnosis and treatment of vascular adverse events with hyaluronic acid fillers. Journal of the American Academy of Dermatology 2019 [Epub ahead of print].
- Delorenzi C. New High Dose Pulsed Hyaluronidase Protocol for Hyaluronic Acid Filler Vascular Adverse Events. *Aesthet Surg J* 2017;**37**(7):814-25.

AESTHETIC FOCUS

AUTHORS



Jani van Loghem,

Aesthetic Physician; Founder and Medical Director, UMA Institute; Amsterdam, The Netherlands. Former Vice Chairman, Dutch Society of Aesthetic Medicine; Head of Department of Aesthetic Medicine, European College of Aesthetic Medicine and Surgery (ECAMS).



Pieter Siebenga,

Aesthetic Physician specialising in injectables, laser and chemical peelings; responsible for the conduct of all clinical research at UMA, as well as the theoretical content of the UMA Academy.



Job Thuis,

Aesthetic Physician, UMA Institute, Amsterdam, The Netherlands; Aesthetic Clinical Educator for Merz Benelux and Key Opinion Leader for Merz Netherlands; and faculty member for ECAMS.

Declaration of competing interests: JVL has been reimbursed by Merz Aesthetics, Allergan, Ipsen and Humanmed for attending several conferences, speaking at several conferences, acting as a consultant and a trainer. JVL is shareholder of UMA Institute and UMA Academy and is lead trainer in the Vascular Complications Management and Retrobulbar Cadaver Injections course, as well as author of the book *Soft Tissue Filler Complications Management in Aesthetic Medicine*, published by UMA Academy and the online webinars on complication management on the UMA Academy website.