INVITED REVIEW

Craniofacial osseointegration

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In year 2007, 30 years have elapsed since the first patient was supplied with a craniofacial osseointegrated implant. The reason for implanting in this patient was a severe conductive hearing loss, which necessitated the use of a bone conduction hearing aid. By utilizing the possibility to transmit sound to the cochlea via direct bone conduction, a new era in audiology was established. Further applications of osseointegration in the craniofacial field is related to the rehabilitation of patients with defects from cancer therapy, malformations, traumatic amputations and burns. Specific fields of osseointegration in this respect are due to possible side effects from radiotherapy and chemotherapy that will affect osseointegration negatively. Other aspects are related to osseointegration in children. This review will focus on the knowledge gained during the first 30 years of craniofacial osseointegration.

Keywords: osseointegration; craniofacial; bone-anchored hearing aid; prosthetics; cancer; malformations

Introduction

Professor Per-Ingvar Brånemark in 1977 coined the term ‘osseointegration’. Although the definition has varied slightly during the years, principally it means ‘a direct structural and functional connection between ordered’, ‘living bone and the surface of a load-carrying implant’ (Brånemark, 1985). Osseointegration was originally used to anchor dental prostheses or bridges in the jaws to replace lost teeth (Brånemark et al., 1969). Since 1965, the concept of osseointegration has profoundly and definitely changed the fundamentals of odontology (Brånemark et al., 1977; Adell et al., 1981; Albrektsson and Wennberg, 2005). It is expected that in the year 2007, more than 8 million osseointegrated implants will be installed in more than 3 million patients for rehabilitation of total or partial edentulousness. Likewise, it has changed the fundamentals of those specialities working outside the oral cavity. These include otolaryngology, head and neck surgery, maxillofacial surgery, plastic surgery, orthopedic-, and hand surgery. The number of patients rehabilitated with these parts of the concept is fewer, but it is estimated that more than 90 000 implants have been installed in more than 45 000 patients until year 2007. This review will focus on the rehabilitation of craniofacial defects – outside the oral cavity – utilizing the osseointegration concept.

Historical aspects

After thorough laboratory work in the 1950s, Brånemark installed the first osseointegrated implant in a human in 1965. It was used for the rehabilitation of an edentulous patient. During a 12-year period, further edentulous patients were included in clinical studies, and in 1977, after examination, the concept of osseointegration was accepted by the Swedish Medical Authorities. The very same year, the first patient had the first extraoral osseointegrated implant installed at our department. This patient was an elderly male with problems of hearing. He was earlier utilizing a bone conduction hearing aid. This aid functions by pressing the vibrator via a steel band to the skull, and it is known to cause pain due to pressure on the skin when using the aid for longer periods. In 1979, a patient with an ear defect caused by cancer was operated upon and supplied ear prosthesis in our department. This was followed by osseointegration in other regions of the craniofacial skeleton as the orbit, nose, and midface during the following years. By 1986, we had altogether operated 100 patients for different craniofacial defects, and then organized the first international workshop on craniofacial osseointegration. The concept for anchoring craniofacial prostheses on osseointegrated implants was accepted by the Food and Drug Administration (FDA) in 1985 and FDA accepted the concept to anchor hearing aids on implants in adults in 1995 and in children 1998.

In 1985, osseointegrated implants were used also in hand surgery to replace defect finger joints (Hagert et al., 1985), and in the beginning of the 1990s the concept was used for orthopedic reasons for rehabilitation of
amputees’ fingers, hands, arms and legs (Brånemark et al., 2001).

**Developmental steps**

To be able to develop a craniofacial osseointegration concept that worked in the clinic, certain steps had to be undertaken. Different kinds of implants were developed from those used in the oral cavity. These were made from commercially pure (C.p.) titanium and were generally shorter or 3–5 mm long, threaded and with the same machined surface as the oral implants. It was further found important to attach a flange in the coronal part of the fixture. The reason for this was the belief that after a longitudinally directed trauma to the implant the flange would prevent it to be pushed into the brain. This has also been shown to be a safe security measure, as several trauma cases have occurred, but only a minority has caused fractures of the skull bone, and none has caused a severe damage (Tjellström, 1989).

The thickness of the skull bone determines the length of the fixtures. Adult temporal bone is usually 4 mm thick in the temporal line, which is also the length of the most commonly used fixtures. In the frontal bone, zygoma and maxilla, longer fixtures are possible to install. With the aim to obtain bicortical anchorage, the fixture should be as long as the bone thickness. If the bone is much thinner, sometimes only 1–3 mm, some patients who have been irradiated or who sustained burns will have an altered texture of bone that will reduce the bone incorporation. A very rough surface will result in rapid integration whereas a smooth surface will result in poor integration. The original Bra¸nemark implant was machined and hence had a relatively smooth surface. It is today known that a very smooth surface will result in poor integration, but with minor resorption. A very rough surface will result in rapid integration, but secondary inflammation and resorption that can jeopardize integration later on (Albrektsson and Wennergren, 2004).

The bone bed into which the implant is installed is of importance. Thus, there is a difference related to if the implant is installed in a child with relatively soft and immature bone, compared with an adult. The very old person, with osteoporosis will integrate the implant to a lesser degree, and implants have been lost to a higher proportion among these (Drinias et al., 2007). Patients who have been irradiated or who sustained burns will have an altered texture of bone that will reduce the capacity to integrate implants (Granström et al., 1993a).

The surgical technique is of major importance. It is necessary that surgery should be non-traumatic, without causing the drilling temperature to rise by more than a few degrees. The fixture should only be handled by titanium instruments and never touched by the gloved hand. It is further important that the surgical field should be protected from fibers, powder and other substances that might hinder osseointegration (Tjellström et al., 1983).

The load of the implant should preferably be in the longitudinal direction. Thus it is important to avoid rotational or cantilever forces, once the implant has
integrated. If forces are distributed in the longitudinal direction, even very high loads can be withstood by the implant during many years of function (Tjellström, 1989).

Factors of importance to lose osseointegration
Loss of integration can be obtained by overload, torsion forces or direct trauma to the implant. Overload might be the result of long extensions of the bar construction or by misplaced implants. Minor torsion forces might induce microfractures in the bone that with time can rupture the delicate integration. Lack of adequate counterforces during tightening of the abutment, might result in immediate implant loss. Tightening of individual magnets used as retention without counterhold might as well cause implant failures (Tjellström, 1989).

The quantity of bone, and also the quality of bone might determine the forces necessary to lose an implant. The most well-known cause for implant failure, is the bone that has been irradiated as part of cancer treatment. Chemotherapy also affects implant survival to a similar degree as radiotherapy. Other factors that might affect implant survival are osteoporosis, steroid medication and diabetes mellitus (Granström, 2005).

Preoperative planning, patient selection and contraindications
To obtain optimum rehabilitation results, it is important to use a team approach (Van Oort et al, 1994; Schwipper and Tilkorn, 2000; Wolfaardt et al, 2005). Each team can have an individual setting, but it generally comprises a surgeon (ENT, oral, maxillofacial or plastic surgeon), a prosthodontist or anaplastologist and a nurse as coordinator. It is valuable to have a radiologist and often other specialists as speech pathologist, audiologist, etc., whenever needed.

Planning craniofacial osseointegration is a multifactorial process and requires tailoring for each patient. An adequate evaluation of the patient’s general health status including psychological profiling prior to surgery is important. A computed tomography (CT) scan or other radiographic evaluation of the bone quantity and quality is essential. Hemoglobin count and other necessary laboratory data are gathered. If the patient has been able to meet another patient with a similar defect, this is helpful for the understanding of the planned rehabilitation. At the performance of surgery, it is essential that the prosthodontist/anaplastologist gives instructions as to the ideal position, number and angulations of implants. CT scan recordings can be analyzed and manipulated in an information technology implant planning application (Wolfaardt et al, 2003). Implant planning software allows bone volumes and densities to be assessed. Implant installation can be simulated, depth of soft tissues overlying the area can be assessed and emergence profiles be considered. A more sophisticated approach to craniofacial implant planning has been described (Watson et al, 1993). This technique makes use of overlaying laser scanning of soft tissues that is digitally overlaid on CT scanning and then CNC milling is used to locate the desired implant positions.

Rapid prototyping with stereolithography has been used in patients with defects of the craniofacial region. The rendered acrylic model of the region assists in the positioning of implants (Wolfaardt et al, 2005).

The indications for implant-based surgery may vary considerably. It is important to judge the alternatives to the planned rehabilitation. The motivation of the patient may likewise vary considerably. It is also important to find out if the patient has realistic expectations on what is possible to obtain. The age of the patient in relation to maturity, and the influence of parents or other relatives must be judged before starting the rehabilitation. Previous surgery in the field of operation and combined radiotherapy/chemotherapy may jeopardize implant installation.

Contraindications to craniofacial osseointegration may be psychiatric disease, alcohol and drug abuse, senility or other disease making the patient unable to take care of implant care. Patients not planning to participate in follow-up programs may also be contraindicated. It is imperative that the patient be prepared to take care of the implant in the followup time.

Any condition that influences the bone to remodel may influence the process of osseointegration. The literature cites numerous potential local and systemic factors that may negatively affect craniofacial osseointegration. It appears that smoking reduces implant survival, although well-designed studies are lacking. The literature on smoking and implant survival addresses dental implants and not craniofacial osseointegrated implants (Esposito et al, 1998).

Equipment and tools
The original craniofacial implant was introduced by Nobelpharma, later Nobel Biocare, Gothenburg, Sweden. This implant system, the Brånemark osseointegration system® was developed from the oral concept utilizing the same drilling equipment, implant surface and drill machine. This implant system has remained relatively unchanged during the years. Only minor alterations of the implants have been undertaken as changed angulations of the abutment for the BAHA. Different angulated abutments – console abutments – have also been developed for use in the orbit. A new self tapping fixture is presently introduced in the market.

There are today several other systems that use other surfaces of the fixture (Conexão®, Otorix®, Straumann®, ITI®), but they all are principally based on the original implant, and hence the same drilling equipment can be used also for these. Surfaces other than the original machined are as well available today.

Surgery for craniofacial osseointegration
Surgery is generally performed under local anesthesia with appropriate premedication. General anesthesia is recommended in children and in persons in whom the drilling procedure will be extended. The surgical installation of screw-shaped osseointegrated implants is well described in the literature (Tjellström, 1985, 1989). The original surgical technique used a two-stage approach with 3–4 months’ healing between the stages of surgery.
Later, a one-stage approach in the mastoid was described (Tjellström and Granström, 1995). The two-stage approach continues to be advocated for pediatric patients, the orbit, midface applications and in patients who have been irradiated (Granström et al, 1993a, 2001; Granström, 2005). Principally, two drilling techniques are used when installing extraoral fixture. During the first technique high speed (2000 r.p.m.) drills are used to make a hole in the bone at the decided place for the fixture. By using sharp drills and profound irrigation, the temperature is kept low, and the bone cells will remain alive. After the site of the implant has been prepared, low speed drilling (15 r.p.m.) is taking place, and the width for the fixture is finally decided and the fixture installed. Today, most often a self-tapping and the width for the fixture is finally decided and the fixture installed. Today, most often a self-tapping implant is used whenever possible. Threading might be necessary in a very hard and brittle bone.

In the adult non-irradiated patient, the attachment of the abutment can proceed immediately after implant installation. In children and irradiated patients, surgery is still conducted in two stages with a healing time of 3–6 months in between. Reduction in the subcutaneous tissue then takes place. This is one of the most important steps, as a thick skin will result in more adverse skin reactions (Reyes et al, 2000). The subcutaneous tissue is removed at a long distance away from the implant to allow a smooth and thin lining of the skin immediately close to the abutment. After surgery, a healing cap with an ointment-soaked gauze rapped around the abutment is used to prevent postoperative bleeding and infection, and to keep the skin pressed to the bone surface during the initial healing period.

**Postoperative treatment**

Most surgeries at our department have been performed under local anesthetics and as day-care surgery. Thus, the patient usually goes home the same day of surgery. The surgical dressing can be removed on the day after surgery while the healing cap and soaked gauze kept for a week. The patient then comes back to the outpatient clinic after a week when the stitches are removed. After the skin and soft tissue has healed, the fitting procedure for the BAE or BAHA can begin.

**Self-care**

It is important that the patient is correctly instructed on how to take care of the skin-penetration site. Debris will invariably collect around the neck of the abutment. This debris is a mixture of keratinocytes and evaporation and should be removed every day with a soft toothbrush or cotton tip, soap and water. If the patient can handle the skin penetration adequately, a long-term reaction free skin is possible to obtain. The patient should be included in a self-care program with daily cleanings, using soap and water, dental floss and soft toothbrush in specific areas. The patient should also attend the clinic for regular follow-up visits, three to four times the first year and twice a year during the following years. At the visits the clinical scores are registered, cleaning is checked and reinstruction given when necessary.

**Applications**

**General indications**

**The bone-anchored hearing aid**

The bone-anchored hearing aid was developed to replace standard bone conduction hearing aids. Bone conduction aids work by pressing the vibrator to the skull via the skin. Better sound transmission is obtained when the steel band presses the vibrator hard on the skin. This is, however, often painful to the patient after a certain time of use. The BAHA functions by a direct sound transmission through the bone, via a permanent skin penetration. Thus the painful pressing of the skin is avoided, and also a better sound transmission is obtained (Tjellström and Häkansson, 1995). Several different types of the BAHA have been developed during the years (Snik et al, 2005). The BAHA Compact is the smallest device, suitable with those patients, often children with the best cochlear function. BAHA Classic is the standard ear level device, and will be gradually replaced by BAHA Divino, which is the first digital BAHA developed. The body-worn device BAHA Cordelle has the strongest output and is used by patients with the poorest cochlear reserve.

Audiological prerequisites for the BAHA is a maximal recommended bone conduction threshold of 45 dB hearing level (HL) taken as an average of 0.5, 1, 2, and 3 kHz for the ear level device. For the body-worn device, a maximal recommended bone conduction threshold of 60 dB (HL) take as an average of 0.5, 1, 2, and 3 kHz.

Indication for BAHA is any patient needing a bone conductor. The most common reason for supplying a patient with BAHA is chronic ear disease, when a standard air conduction aid occludes the ear canal, and the ear starts to drain (Snik et al, 2005). The second most common reason is bilateral ear canal atresia. Conductive hearing loss in only hearing ear is also an indication, as is external otitis, which prevents the patient from using standard air conduction aids.

The prescription of bilateral BAHA is recommended in certain cases. This can be persons with specifically demanding occupations as teachers and musicians, with high demands on sound localization. The amplification from double BAHAs is in the range 5 dB (Priwin et al, 2004). Patients with unilateral ear canal atresia, can sometimes gain from a BAHA on the malformed side. This goes especially for children (Priwin et al, 2007). The latest indication for BAHA is single-sided deafness, in which patients can obtain some directional hearing from the shadow side (Snik et al, 2005).

**Bone-anchored epistheses**

**Ear prosthetics.** Cancer surgery, malformations, traumatic amputations or burns can cause ear defects. Alternatives to rehabilitate the patient may be to disclose the defect with a natural hair styling. This is fully satisfactory to some patients. Autogenous reconstruction has become a realistic procedure in the last decade (Firmin, 1998). Several plastic surgeons can make very natural looking autogenously reconstructed ears, and...
this should therefore be the principal option. If the reconstruction fails, it can easily be removed, and an implant-retained episthesis be an alternative (Wilkes and Wolfaardt, 1994). Radiation therapy, scarring due to trauma or burns may compromise the local tissue so that autogenous reconstruction may be contraindicated and craniofacial osseointegration is the only realistic treatment option. Anatomically, the lower half of the ear is the most challenging to reconstruct (Wolfaardt et al., 2003). In those cases where a prosthesis is indicated, the anchorage could be by glue, undercuts, spectacles or by implants. Bone-anchored ear episthesis has an advantage over the earlier mentioned retention systems in that it is a safe, easy and secure anchoring method (Granström et al., 1993b). Where attempts of autogenous reconstruction have failed, craniofacial osseointegration provides a valuable salvage option. A particularly controversial aspect of treatment selection in ear reconstruction is in the pediatric patient with microtia. It is possible to place osseointegrated implants in children, but the question remains as to what is appropriate. If an autogenous reconstruction fails, the craniofacial osseointegration portion remains. However, if the patient rejects an implant-based prosthesis, an autogenous option might not be available to the patient. With the installation of implants, the microtia area is scarred and this may limit the potential for satisfactory autogenous reconstruction if indicated later. Consequently, it is important to ensure that the patient and parents are well informed of their options before treatment proceeds.

**Orbit prosthetics.** Defects of the orbit can appear because of cancer surgery, after certain eye diseases, traumatic ablatio and burns. Alternatives to disclose the defect can be to cover the defect with a patch or supply the patient with prosthesis anchored on spectacles, undercuts or glue. Autogenous reconstruction may be limited when the contents of the orbit have been exenterated or severely anatomically disrupted. Skin flaps may be used to provide coverage of the orbit but provide poor aesthetic results. A prosthesis based on osseointegrated implants is the best option for several reasons (Wilkes and Wolfaardt, 2000). Surgery is simple and straightforward, and the implant is easily removed if the patient changes his/her mind. An important aspect is that it is easy to inspect the tumor cavity if the patient wears a prosthesis. If a skin flap covers the defect, recurrent tumor will be detected later.

**Nose prosthetics.** Nose defects most often occur after cancer surgery, traumatic amputations (animal bites) and burns. Alternatives to rehabilitate the patient can be to cover the defect with a patch, or prosthesis anchored on spectacles, undercuts, glue or osseointegrated implants. Autogenous reconstruction of the nose is possible to make with high esthetical results, and might therefore be the first option for the patient (Weiss et al., 1998). In common with aspects of ear reconstruction, if the patient’s medical status precludes surgery, residual tumor is present, there are no suitable donor sites, the patient will not tolerate the donor sites or by the patient’s choice, autogenous reconstruction may be precluded. In these situations, craniofacial osseointegration becomes the treatment of choice. Where there has been severe loss of facial contour this may be best reconstructed with an implant retained nasal prosthesis. If one, however, decides to supply the patient with an implant-based episthesis for reasons of cancer observation, secondary autogenous reconstruction is still possible years later. That is another advantage with osseointegration, in that it is a reversible procedure if the patient changes his/her mind.

**Midface prosthetics.** Midface defects are often a result of cancer surgery, gunshot wounds, burns, and in some occasions caused by malformations, e.g. non-reconstructed lip, jaw and palate clefts. In limited defects of the midface, autogenous reconstruction, combining different grafting techniques might be the first choice. Where the defect extends to the oral cavity or orbit, where an autogenous reconstruction is not possible, osseointegrated implants may offer the most appropriate treatment option (Eckert and Desjardin, 1998; Harris et al., 1996; Schwipper and Tilkorn, 2000). In a number of cases, combined grafting and osseointegrated implants for prosthetics is favorable.

**Major facial prosthetics.** Major facial defects are often a result of cancer surgery, gunshot wounds or burns. The first option is often autogenous reconstruction, combining different grafting techniques. In those instances where orbit or nose is deficient, a prosthetic solution can be combined with autogenous reconstruction. Here, episthesis anchored on spectacles, undercuts, glue or implants is an option. In several cases, the combined grafting and osseointegrated implants for prosthetics is favorable.

**Other areas in the craniofacial region**

Where significant areas of hair loss have occurred, osseointegrated implants have been used to retain a hairpiece (Weischer and Mohr, 2000, Wolfaardt et al., 2005). While this is not a widely used application of craniofacial osseointegration, it may provide an option where wearing of a conventional wig, hairpiece, tissue expansion or hair transplantation is not possible.

**Grafting as part of procedure**

Especially in cancer patients, the defects from cancer surgery may be extensive, and there might be limited bone left, available for implant installation. The tissue may be further compromised if the patient has received irradiation and chemotherapy as part of cancer treatment. In these cases, grafting of the bone to the implant site may be an option. There are several techniques used to bring the bone, soft tissue and skin to the craniofacial region (Urken et al., 1989; Triplett et al., 2000). The specific technique utilized must often be adapted to the patient’s specific defect. Placement of craniofacial implants at the time of surgical reconstruction has some advantages, particularly when the implants help to stabilize the graft in relation to the host bone. As fewer surgical procedures
are needed, patient morbidity is potentially reduced and the process may be cost-effective. Secondary implant insertion does, however, allow more precise planning of the optimum implant position and angulation (Tripelett et al., 2000). Some authors do not support the use of primary installation of craniofacial implants in grafts, as the risk for misplacement is obvious. It is further reported a higher risk of loosening implants, risk for osteoradionecrosis and risk for survival of graft in those cases (Schmelzeisen et al., 1996; Schoen et al., 2001). The prognosis for implants placed in a vascularized bone graft is generally better than those placed in non-vascularized grafts, whether in a primary or secondary implant placement (Flood and Russell, 1998).

Results and follow-up
By the end of 2006, a total of 1453 patients had been rehabilitated by the osseointegration concept at our department. The most common reason for implantation was the BAHA (869 patients), followed by epistheses for ear reconstruction (342), orbit (116), different facial epistheses (42) and maxillary or mandibular defects (84). Of the reasons for rehabilitating a patient with an episthesi, cancer was the most common (422) followed by malformations (128), burns and trauma (37). More than 3000 osseointegrated implants have been installed in the craniofacial region without any serious complications until today. During 20 years of follow-up, implant failures have been approximately 10% (Tjellström and Granström, 1994; Reyes et al., 2000). There are certain factors that determine the implant failure rate. The temporal bone shows the lowest implant failure rate or 8% over a 10-year period. This is a relatively compact bone with properties similar to that of the mandible. In this bone, it is possible to detect an age-dependent difference in failure rate (Drinias et al., 2007). Thus the oldest patients showed the highest failure rates, probably as a result of increasing osteoporosis. In the other craniofacial regions, the highest implant failure rates were seen in the frontal bone (50%) and zygoma (20%) (Granström et al., 1994). This has been found related to these bones being irradiated more often than the other regions (Granström, 2005).

The skin penetration
The skin penetration site is the single factor that has caused craniofacial osseointegration the most significant clinical problems (Abu-Serriah et al., 2001). Despite extensive subcutaneous reduction during surgery, some patients will experience a red and moist skin, and sometimes granulation tissue forms around the abutment. Compared with oral osseointegration, some differences are obvious. The gingiva is constructed to have a mucosal penetration. Saliva and the cleaning ability of the oral tongue contribute to good condition (Steffick et al., 1991; MacKenzie and Tonetti, 1995). Nevertheless, most patients can obtain and maintain a reaction free skin penetration for many years (Tjellström et al., 1983, 1985). To be able to control the adverse reactions, a skin condition scoring system was developed (Holgers et al., 1987) where 0 is no reaction; 1: reddish; 2: red and moist; 3: granulation tissue; 4: skin infection to such a degree that the abutment has to be removed. The scores vary depending on how many implants have been inserted, the region in which the implants were inserted, the retention system used and the age of the patient. Thus 92.5% of BAHA users had a reaction free skin (score 0), 91.1% of orbital prosthesis users and 89.3% of auricular prosthesis users were similarly free of complications (Holgers et al., 1992). Young children tend to have low reaction scores during childhood when the parents take care of the implants. In adolescence, however, the adverse reactions tend to increase as the care of implants is neglected (Granström et al., 1993b). Of those patients experiencing adverse reactions, 15% contribute to more than 70% of all adverse reactions and the reason for the skin reactions can be explained by insufficient home care (Tjellström, 1989). Over longer times, the number of adverse skin reactions decline as an indication that the patients learn how to take care of the skin-penetration site (Reyes et al., 2000). Some authors have suggested that the subcutaneous tissue in the peri-abutment region should be thinned or a split thickness skin graft should be placed so as to limit the tissue movement (Albrektsson et al., 1987; Tjellström, 1990; Tolman and Desjardins, 1991; Westin et al., 1999). Limiting movement will, theoretically, decrease the shearing forces at the soft tissue–implant abutment interface, and better maintain a barrier against bacterial ingrowth. However, some may argue that trimming of the subcutaneous tissues or split skin grafting can result in scarring and so compromise the vascularity of the peri-implant soft tissue, and increase the potential for adverse skin reactions. Surgical manipulation of the subcutaneous tissue may also alter connective tissue–epithelial interactions and undermine the integrity of the implant–soft tissue interface. The importance of healthy underlying the connective tissues has been well recognized, but it is not clear just how thick the peri-abutment skin should be (Abu-Serriah et al., 2003).

Maintaining skin problems
When the patient attends the clinic with a grade 1 reaction, we have intensified the cleaning at the follow-up visit, re-instructed the patient and given him an earlier time for the next check-up visit. When grade 2 reactions occur we have done as for grade 1 and furthermore prescribed local ointment (Terracortril + Polymyxin B®; Pfizer, Dublin, UK), which contains antibiotics, antimycotics and steroids. This is applied 2–3 times a day for the next week and the patient returns to the clinic for our inspection. When grade 3 reactions occur we have done as for grade 2 but also wrapped a Terracortril and Polymyxin B-soaked tampon around the abutments with changes for a week. An alternative could be to remove the bar and put on the healing caps around which the tampon is wrapped. If the tissue reaction is not under control, surgical removal of the granulation tissue followed by new skin grafting is suggested. A grade 4 reaction means that all attempts to correct a grade 3 reaction have failed. We then remove the abutment and leave the skin for
secondary healing. If necessary a new abutment could be reconsidered later on.

Certain skin conditions need specific attention. Acne vulgaris can create problems in the skin penetration area during adolescence, and sometimes afterwards. It can generally be handled by local cleaning or by drugs prescribed for the disease. Seborrhoic eczema can cause local problems, especially in the temporal region. Local treatment with steroids can be helpful as grafting of non-eczematous skin. Psoriasis can be treated in a similar way, but may need proper planning to avoid primary installation in the lesion. Patients with diabetes mellitus often have a reduced healing rate, and skin necrosis is more common. Secondary healing is often uneventful in any case. Keloid formation can be abundant, especially in burns patients. Local excision combined with injection of steroids might help reduce the problem.

Irretractable skin problems can be due also to loose fixtures or abutments. These should be checked and retightened upon the patient follow-up visits. Misplaced implants, with too narrow positioning might also affect the skin condition, as it can be very difficult for the patient to clean (Tjellström, 1989; Reyes et al, 2000).

Osseointegration in irradiated patients
This topic has caused some controversy during the years (Donoff, 2006; Granström, 2006). The fundamental question is whether radiotherapy causes negative effects on osseointegration or not. Increased knowledge today shows that a number of factors affect implant survival in the irradiated bone. These include radiotherapy given before or after implant installation (Granström et al, 1993a,c; Granström, 2005). The combined pre- and postoperative irradiation is particularly damaging to implant integration (Granström and Tjellström, 1997). Factors that affect implant survival, such as irradiation dose and fractionation, chemotherapy, time from radiotherapy to implant surgery, and also other factors such as fixture length and prosthetic retention affect the results. Implant survival also depends on the length of time that one follows the patient. (Granström, 2005). Despite the fact that implant survival might be affected by radiotherapy, the benefits the patient can gain from receiving implants are so high that it is recommended. However, it is also stressed that cancer patients who are supposed to receive osseointegrated implants, should be treated at institutions well used to handling cancer patients. Risk for induction of osteoradionecrosis is always present, and it is of the utmost importance that such side effects are avoided. Adjunctive use of hyperbaric oxygen therapy can reduce that risk (Granström et al, 1994, 1999).

Osseointegration in children
In oral osseointegration it is well known that installation of osseointegrated implants before finished growth of the jaws results in unfavorable implant positioning (Ödman et al, 1991). However, in extraoral osseointegration children may also need to be implanted. These include especially syndromic children with bilateral ear malformations. Not knowing the lowest age possible to implant children, a gradual lowering of age was used to gain knowledge of factors of importance for obtaining stable implants in this patient category. The thickness of the parietal and temporal bones is a limiting factor for implant installation. Measuring the bone thickness during surgery shows that a lower age of approximately 4 years is necessary to have a reasonably thick bone for a 4-mm fixture to be installed (Priwin and Granström, 2005). In children who do not have this required thickness of the skull bone, it is still possible to install fixtures with bone augmentation by use of semipermeable membranes (Granström and Tjellström, 1999). An increased bone thickness of 1–2 mm is possible to obtain during a healing period between the first and the second stage surgery of 4–6 months (Granström and Tjellström, 1999). With the use of this technique it has been possible to implant children down to an age of 1 year without significantly higher implant failures.

Developmental projects
There are several unresolved problems related to craniofacial osseointegration, which are subject to research at our institution. The skin penetration is the most prominent clinical problem in many clinics utilizing craniofacial osseointegration. Despite surgical techniques leading to a thin, non-mobile skin penetration, several patients develop gradually thicker skin with related problems due to cleaning difficulties. This might lead to dermatitis and possible risk for implant failure with time. We are seeking an increased understanding of the biological behavior of the skin. Our hypothesis is that several of the factors of importance for inflammation of the periodontium is present in the skin as well. Thus, introducing methods of determining which inflammatory systems are active might help to understand why dermatitis occurs at the penetration site. We are looking further for alternative medical and surgical treatment options.

Irradiation of the bone bed is a topic of further study. Knowingly that implant failures are higher in the previously irradiated bone, new approaches to improve osseointegration are necessary. This project is a multicentre study performing a randomized, controlled study comparing normobaric air to hyperbaric oxygen. Interested readers can enroll patients on the website http://www.oxynet.com. In future studies pharmacological effects, e.g. from growth factors in the compromised tissues are to be studied. New implant materials are tested in several controlled studies. These include implants of different sizes and shapes, with modified surfaces. These studies also utilize non-interventional measurement of osseointegration as resonance frequency analysis and different radiological techniques. Computer-aided techniques to better control fixture placement will be utilized. Osseointegration in children is subject of further studies with the aim to determine the optimum age for implantation. Several of the children have reduced bone at the implant site necessitating bone augmentation. The future growth of the temporal bone is a factor that needs to be controlled. With time, the fixtures will be buried in an inferior position, which might need further surgery to adjust. In a research project, a superficially positioned implant will be inserted.
already from the beginning to compensate for future growth. Two projects studying implantable BAHAs, and BAHAs for single-sided deafness are conducted. These projects aim at rehabilitating persons with hearing handicaps that are not possible to treat today.

Aspects for the future
The development of craniofacial osseointegration will continue during the next decades and further. New macrostructures that can affect bone interactions will be developed. These could include different angulations of the threads, addition of microgrooves and changed anatomy of the implant neck and additional undercuts in the fixture as described for oral implants (Miranda-Burgos, 2006).

New surfaces that will stimulate bone formation and add years to successful osseointegration will be developed. New surgical techniques that are faster and simpler both for the patient and surgeon will be developed. Growth factors, stem cells and other pharmacological drugs will help us to improve osseointegration in compromised tissues (Thor, 2006). We will have help from non-intervention evaluation of the implants to understand better the biology of osseointegration. New prosthetic materials that help us rehabilitate patients with severe craniofacial defects will be developed.

It will be possible to build together form and function combining osseointegration with microelectronics, e.g. as described by Klein et al., 1999). And more is to come that we cannot tell today.

References