

IMMEDIATE LOADING IMPLANTS: REVIEW OF THE CRITICAL ASPECTS

L. TETTAMANTI¹, C. ANDRISANI², M. ANDREASI BASSI³, R. VINCI⁴,
J. SILVESTRE-RANGIL⁵, A. TAGLIABUE¹

¹ Department of Medicine and Surgery, University of Insubria, Varese, Italy

² Private Practice in Matera, Matera, Italy

³ Private Practice in Rome, Rome, Italy

⁴ Oral Surgery, Università Vita-Salute San Raffaele, Milan, Italy

⁵ Department of Estomatology, University of Valencia, Valencia, Spain

SUMMARY

Purpose. Modern dentistry have witnessed, a rapid and continuing evolution. Concerning the implant-rehabilitation protocols, they have been redefined in order to satisfy patient's increasing expectations in terms of comfort, aesthetic and shorter treatment period. The purpose of this review is to explore the concept of implant immediate loading and the indications for clinical practice. All the critical aspects that could influence the outcomes of this treatment will also be considered.

Materials and methods. Three protocols for implant load timing have been classified: immediate loading implants (ILI); early loading implants (ELI); and conventional loading implants (CLI). Two subclassifications point out the different loading modality: 1) Occlusal loading or Non-Occlusal loading, 2) Direct loading or Progressive loading. Micromovements have been considered, since the start of implant dentistry, one of the main risk for the success of osseointegration. The determinant and most accessible parameter to assess the primary stability is the implant insertion torque value. To achieve the necessary torque value to perform immediate loading, it is therefore important to evaluate the bone density at the implant site. Computerized tomography (CT) has been regarded as the best radiographic method to evaluate the residual bone.

Results. The clinical success of this technique is highly dependent on many factors: patient selection, bone quality and quantity, implant number and design, implant primary stability, occlusal loading and clinician's surgical ability. Among these, implant primary stability is undoubtedly the most important.

Conclusion. Studies on ILI show that successful outcome can be expected, if the previous criteria are fulfilled. It seems that ILI demonstrate a greater risk for implant failure when compared to CLI, although the survival rates were high for both the procedures. The use of different surgical procedures, type of prostheses, loading times and have very different study designs. This lack of homogeneity limits the relevance of the conclusions that can be drawn.

Key words: immediate loading implants, implant stability, implant design.

Modern dentistry has witnessed, over the last decades, a rapid and continuing evolution of techniques in different fields (1-44, 122-125). Concerning the implant-rehabilitation protocols, they have been redefined over the years, as a result of new knowledges in implant surgery and in order to satisfy patient's increasing expectations in terms of comfort, aesthetic and

shorter treatment period.

Since Branemark introduced the osseointegration system in 1977 (45), new protocols have been proposed regarding the prosthetic-load timing, up to the immediate implant loading.

Classic protocols propose that implants should receive no loading during the osseointegration period, usually 3 to 4 months in the mandible

and 6 to 8 months in the maxilla (46-48). Updated protocols have shortened the healing period, so that implants could be loaded early and even immediately, before osseointegration is completely obtained.

The purpose of this review is to explore the concept of implant immediate loading and the indications for clinical practice. All the critical aspects that could influence the outcomes of this treatment will also be considered.

■ Implant loading time protocols

Esposito et al. (49) have defined 3 protocols for implant load timing: immediate loading implants (ILI), within 1 week from implant placement; early loading implants (ELI), between 1 week and 2 months; and conventional loading implants (CLI), after 2 months from implant placement.

Two subclassifications point out the different loading modality: 1) Occlusal loading or Non-Occlusal loading, 2) Direct loading or Progressive loading.

This Cochrane systematic review (49) concludes that there is no convincing evidence of a clinically important difference in prosthesis failure, implant failure, or bone loss associated with different loading times of implants.

The results of a meta-analysis by Enríquez-Sacristán et al. (50) report that ILI, ELI and CLI share similar success and survival rate.

A recent meta-analysis, by Sanz-Sánchez et al. (51), together with a recent study by Zhu et al. (52), shows that ILI demonstrate a greater risk for implant failure when compared to CLI, although the survival rates were high for both the procedures.

Moreover ILI, ELI and CLI were found not significantly different in terms of the associated marginal bone loss, changes in implant stability, and health status of the peri-implant tissues, which indicated that these loading protocols

behaved similarly once osseointegration occurred (52).

■ Primary stability

Micromovements have been considered, since the start of implant dentistry, one of the main risk for the success of osseointegration (47). It has been proved that if the micromovements range results to be over 150 μm , this could jeopardize the osseointegration process. This excessive micromotion results to be directly implicated in the formation of the implant fibrous encapsulation (53, 54). The literature suggests that there is a critical threshold of micromotion above which fibrous encapsulation prevails over osseointegration. This critical level, however, was not zero micromotion as generally interpreted. Instead, the tolerated micromotion threshold was found to lie somewhere between 50 and 150 microns (54, 126).

In this tolerated micromovements range, an early load on the implant surface could even stimulate the newly formed bone to remodel, accelerating the osseointegration process.

That being said, all the studies in literature agree that achieving good implant primary stability is key condition to ILI success (55).

Primary implant stability is influenced by many factors including local bone quality and quantity, implant macro-design and surgical technique (56, 127-132).

■ Implant primary stability evaluation

The determinant and most accessible parameter to assess the primary stability is the implant insertion torque value.

Torque values ranging from 30 to 40 Ncm and higher have been usually chosen as thresholds for immediate loading (57, 58). That torque minimum level is important both to assure the

osseointegration process and to give enough engaging strength to the implant-abutment connections, via the fixation screw. Nonetheless, some studies assess that also ILI placed in a weak bone with a final torque ≥ 20 Ncm have an equally successful prognosis as the CLI (59).

Furthermore, if enough implants are placed, ILI can be performed even if not all the implants achieve an adequate stability, thanks to the support of adjacent implants, but the unstable implants should be left unloaded (60).

To measure the implant primary stability, recently it has been developed an Implant Motor (TMM2®, Idievolution) that allows the clinician to measure the bone density during both the preparation of the implant site and the implant insertion.

Two other methods to measure the primary stability are the resonance frequency analysis (RFA) and the Periotest® (PT).

The RFA (Osstell®) is a reliable device that measures the resonance frequency of a transducer attached to the implant body (61). The result of the measurement is the implant stability quotient (ISQ), which reveals the hardness of the implant-bone connection (62). ISQ values greater than 65 have been regarded as most favorable for implant stability, whereas ISQ values below 45 indicate a poor primary stability (63).

The PT indicates implant stability by measuring the time of contact between the instrument's tip and the implant, during repetitive percussions generated by this device.

The lack of well-defined reference values, both for the RFA and for the PTV, and the possibility of some operator-dependent variations in the measurements, make their routine clinical use not efficient (64).

Bone quality and quantity

To achieve the necessary torque value to perform immediate loading, it's therefore impor-

tant to evaluate the bone density at the implant site. Computerized tomography (CT) has been regarded as the best radiographic method to evaluate the residual bone (65).

Several classifications regarding bone density have been proposed. In 1990, Misch (66) proposed a classification based on macroscopic cortical and trabecular bone characteristics: Class I: dense cortical bone; Class II: porous cortical bone; Class III: coarse trabecular bone; Class IV: fine trabecular bone.

When Class III or Class IV bone is present at implant site, the operator can overcome this limitation performing specific surgical techniques and using implant with peculiar macro surfaces.

Surgical techniques

In several studies, Authors introduced different techniques to locally optimize bone density and subsequently improve primary stability, such as 1-2 mm subcrestal implant placement (67, 68), bicorticalization into the nasal or sinus floor whenever possible (69), implant site under preparation (70) and bone condensing technique (71). The implant site under preparation and the bone condensing technique are the most commonly used techniques, performed nearly always when in presence of Class III or Class IV bone: the first consists in the use of a final drill diameter which is smaller than the diameter of the implant (70); with the second technique cancellous bone is pushed aside with bone condensers (osteotomes), thus increasing the density of the implant surrounding bone (71).

Through the use of these procedures, it has been reported high survival rates with ILI (67, 72).

In areas where bone augmentation is needed, CLI should be the first choice (48, 73, 74). Most titles on ILI do not use bone graft or sinus lift procedures. Recent studies report successful use of bone grafts to fill horizontal gaps be-

tween the implant surface and the extraction socket walls and to cover the vestibular dehiscences (60,73).

Implant design and positioning

Regarding the implant macro-design, tapered (root-form) implants were introduced to overcome the poor bone quality and quantity limitations. The goal behind using tapered implant was to exercise a degree of compression of the surrounding bone during the insertion phase, and the decrease of their apical diameter allows to accommodate them in area with small bone volume available, like the labial concavity or between adjacent roots (75).

Implant surface characteristics and diameter have also been shown to influence primary stability: rough implant surfaces make the area of implant-bone contact even more extended (76). Clinical studies have shown that, in cases with a limited bone volume, implants with less than 3 mm diameters can reach sufficient primary stability (77). Wider implants are often used in posterior regions with poor quality bone (70). Single teeth implants demonstrate greater risk of failure, when compared to immediately loaded full arch restorations (51).

To obtain full-arch rehabilitation with ILI, most studies consider 6 implants to be the lowest adequate number to achieve a predictable outcome (36, 37). Malo et al. (25) described a technique to achieve successful results with only 4 implants.

Regarding implant position, all studies give importance to an uniform distribution along the alveolar arch (78); distal implants should be inserted in place of the 2nd premolar or 1st molar, even with a tilted position in order to minimize the need of cantilevers (78). The avoidance of distal cantilevers is considered a success key by many Authors.

Regarding implant length, all Authors prefer using longer implants whenever possible, with

a minimum of 8 mm length, being 13 and 15 mm implants the most frequently used (78). Tilting may enable placement of longer implants in posterior regions (78).

Computer-guided surgery minimize the errors in implant positioning compared to manual or conventional surgical guide implant placement (79), resulting in lesser post-operative morbidity and increased patient satisfaction (80).

Patient selection

Of course, when performing ILI, the patient selection criteria can influence the success of this technique (81).

Most studies in literature propose the following criteria: good general health, edentulous area or teeth with impossible prognosis, adequate bone quality and quantity, absence of acute infection, and primary stability of implants. The exclusion criteria are: systemic disease, immunodeficiencies, head and neck radiotherapy, alcohol or drug abuse, pregnancy, pathologies of the oral mucosa, or lack of cooperation of the patient.

There is no consensus on bruxism or smoking habits (78).

Complications

According to different Authors (82, 83), the ILI protocol more often leads to technical complications. The most common of those were fractures of the prostheses, loosening of the abutment screws and denture contouring adjustments.

The last could be explained by the secondary gingival healing after surgery in the early loading prostheses, which may result in space around the abutments, while relining impressions performed in the conventional loading implants after a period of healing avoided this space (65).

Anyhow, all these complications are solved by adjusting the implants or prostheses without affecting the outcomes of the procedures (78). All together the above mentioned variables are of paramount importance to reduce the risk of peri-implantitis (34, 35, 37, 84-119).

Conclusions

Literature data showed that ILI could represent a reliable and effective protocol to rehabilitate single or multiple missing teeth and offers important advantages for the patient, in terms of function, aesthetics and comfort. However the clinical success of this technique is highly dependent on many factors: patient selection, bone quality and quantity, implant number and design, implant primary stability, occlusal loading and clinician's surgical ability. Among these, implant primary stability is undoubtedly the most important.

Studies on ILI show that successful outcome can be expected, if the previous criteria are fulfilled. It seems that ILI demonstrate a greater risk for implant failure when compared to CLI, although the survival rates were high for both the procedures (120).

Single teeth implants demonstrate greater risk of failure, when compared to immediately loaded full arch restorations (51).

Shimmel M. et al. (121) in a recent review concluded that although all three loading protocols provide high survival rates, ELI and CLI protocols are still better documented than ILI and seem to result in fewer implant failures during the first year.

Studies available use different surgical procedures, type of prostheses, loading times and have very different study designs. This lack of homogeneity limits the relevance of the conclusions that can be drawn.

References

1. Rigo L, Viscioni A, Franco M, Lucchese A, Zollino I, Brunelli G, Carinci F. Overdentures on implants placed in bone augmented with fresh frozen bone. *Minerva Stomatol.* 2011;60:5-14.
2. Carinci F, Brunelli G, Franco M, Viscioni A, Rigo L, Guidi R, Strohmeier L. A retrospective study on 287 implants installed in resorbed maxillae grafted with fresh frozen allogeneous bone. *Clin Implant Dent Relat Res.* 2010;12:91-8.
3. Viscioni A, Rigo L, Franco M, Brunelli G, Avantiaggiato A, Sollazzo V, Carinci F. Reconstruction of severely atrophic jaws using homografts and simultaneous implant placement: a retrospective study. *J Oral Implantol.* 2010;36:131-9.
4. Franco M, Rigo L, Viscione A, et al. CaPO4 blasted implants inserted into iliac crest homologue frozen grafts. *The Journal of oral implantology.* 2009; 35:176-80.
5. Viscioni A, Franco M, Rigo L, Guidi R, Brunelli G, Carinci F. Implants inserted into homografts bearing fixed restorations. *Int J Prosthodont.* 2009; 22:148-54.
6. Franco M, Viscioni A, Rigo L, Guidi R, Zollino I, Avantiaggiato A, Carinci F. Clinical outcome of narrow diameter implants inserted into allografts. *J Appl Oral Sci.* 2009;17:301-6.
7. Viscioni A, Franco M, Rigo L, Guidi R, Spinelli G, Carinci F. Retrospective study of standard-diameter implants inserted into allografts. *J Oral Maxillofac Surg.* 2009;67:387-93.
8. Carinci F, Brunelli G, Zollino H, et al. Mandibles grafted with fresh-frozen bone: An evaluation of implant outcome. *Implant Dentistry.* 2009;18:86-95.
9. Carinci F, Brunelli G, Zollino I, et al. Mandibles grafted with fresh-frozen bone: an evaluation of implant outcome. *Implant Dent.* 2009;18:86-95.
10. Franco M, Tropina E, De Santis B, Viscioni A, Rigo L, Guidi R, Carinci F. A 2-year follow-up study on standard length implants inserted into alveolar bone sites augmented with homografts. *Stomatologija.* 2008; 10:127-32.
11. Danza M, Paracchini L, Carinci F. Tridimensional finite element analysis to detect stress distribution in implants. *Dental Cadmos.* 2012;80:598-602.
12. Danza M, Grecchi F, Zollino I, Casadio C, Carinci F. Spiral implants bearing full-arch rehabilitation: Analysis of clinical outcome. *Journal of Oral Im-*

- plantology. 2011;37:447-55.
13. Danza M, Zollino I, Avantaggiato A, Lucchese, A & Carinci, F. Distance between implants has a potential impact of crestal bone resorption. *Saudi Dental Journal*. 2011;23:129-33.
 14. Carinci F, Danza M. Clinical outcome of implants inserted in piezo split alveolar ridges: A pilot study. In: ed. eds. *Perspectives on Clinical Dentistry*. 2011: 29-30.
 15. Danza M, Zollino I, Guidi R, Carinci F. Computer planned implantology: Analysis of a case series. In: ed. eds. *Perspectives on Clinical Dentistry*. 2011: 287-300.
 16. Danza M, Carinci F. Flapless surgery and immediately loaded implants: a retrospective comparison between implantation with and without computer-assisted planned surgical stent. *Stomatologija*. 2010;12:35-41.
 17. Danza M, Quaranta A, Carinci F, Paracchini L, Pompa G, Voza I. Biomechanical evaluation of dental implants in D1 and D4 bone by Finite Element Analysis. *Minerva stomatologica*. 2010;59: 305-13.
 18. Danza M, Riccardo G, Carinci F. Bone platform switching: a retrospective study on the slope of reverse conical neck. *Quintessence Int*. 2010;41:35-40.
 19. Danza M, Fromovich O, Guidi R, Carinci F. The clinical outcomes of 234 spiral family implants. *J Contemp Dent Pract*. 2009;10:E049-56.
 20. Calvo-Guirado JL, Ortiz-Ruiz AJ, Lopez-Mari L, Delgado-Ruiz R, Mate-Sanchez J, Bravo Gonzalez LA. Immediate maxillary restoration of single-tooth implants using platform switching for crestal bone preservation: a 12-month study. *Int J Oral Maxillofac Implants*. 2009;24:275-81.
 21. Danza M, Guidi R, Carinci F. Comparison Between Implants Inserted Into Piezo Split and Unsplit Alveolar Crests. *Journal of Oral and Maxillofacial Surgery*. 2009;67:2460-65.
 22. Danza M, Scarano A, Zollino I, Carinci F. Evaluation of biological width around implants inserted in native alveolar crest bone. *Journal of Osseointegration*. 2009; 1:73-76.
 23. Danza M, Zollino I, Guidi R, Carinci F. A new device for impression transfer for non-parallel endosseous implants. *Saudi Dental Journal*. 2009;21: 79-81.
 24. Andreasi Bassi M, Lopez MA, Confalone L, Gaudio RM, Lombardo L, Lauritano D. Clinical outcome of a two-piece implant system with an internal hexagonal connection: a prospective study. *J Biol Regul Homeost Agents*. 2016;30:7-12.
 25. Danza M, Guidi R, Carinci F. Spiral family implants inserted in postextraction bone sites. *Implant Dent*. 2009; 18:270-8.
 26. Lucchese A, Carinci F, Saggese V, Lauritano D. Immediate loading versus traditional approach in functional implantology. *European Journal of Inflammation*. 2012;10:55-58.
 27. Traini T, Danza M, Zollino I, et al. Histomorphometric evaluation of an immediately loaded implant retrieved from human mandible after 2 years. *International Journal of Immunopathology and Pharmacology*. 2011;24:31-36.
 28. Scarano A, Murmura G, Carinci F, Lauritano D. Immediately loaded small-diameter dental implants: evaluation of retention, stability and comfort for the edentulous patient. *European Journal of Inflammation*. 2012;10:19-23.
 29. Degidi M, Piattelli A, Carinci F. Clinical outcome of narrow diameter implants: a retrospective study of 510 implants. *J Periodontol*. 2008;79:49-54.
 30. Degidi M, Piattelli A, Iezzi G, Carinci F. Do longer implants improve clinical outcome in immediate loading? *Int J Oral Maxillofac Surg*. 2007;36:1172-6.
 31. Degidi M, Piattelli A, Carinci F. Immediate loaded dental implants: comparison between fixtures inserted in postextractive and healed bone sites. *J Craniofac Surg*. 2007;18:965-71.
 32. Degidi M, Piattelli A, Iezzi G, Carinci F. Retrospective study of 200 immediately loaded implants retaining 50 mandibular overdentures. *Quintessence Int*. 2007;38: 281-8.
 33. Degidi M, Piattelli A, Iezzi G, Carinci F. Immediately loaded short implants: analysis of a case series of 133 implants. *Quintessence Int*. 2007;38:193-201.
 34. Degidi M, Piattelli A, Iezzi G, Carinci F. Wide-diameter implants: Analysis of clinical outcome of 304 fixtures. *Journal of Periodontology*. 2007;78: 52-58.
 35. Degidi M, Piattelli A, Gehrke P, Felice P, Carinci F. Five-year outcome of 111 immediate nonfunctional single restorations. *J Oral Implantol*. 2006;32:277-85.
 36. Degidi M, Piattelli A, Carinci F. Parallel screw cylinder implants: Comparative analysis between immediate loading and two-stage healing of 1005 dental implants with a 2-year follow up. *Clinical Implant Dentistry and Related Research*. 2006;8: 151-60.
 37. Degidi M, Piattelli A, Gehrke P, Carinci F. Clinical

- outcome of 802 immediately loaded 2-stage submerged implants with a new grit-blasted and acid-etched surface: 12-month follow-up. *Int J Oral Maxillofac Implants*. 2006;21:763-8.
38. Degidi M, Piattelli A, Felice P, Carinci F. Immediate functional loading of edentulous maxilla: a 5-year retrospective study of 388 titanium implants. *J Periodontol*. 2005;76:1016-24.
 39. Falisi G, Severino M, Rastelli C, et al. The effects of surgical preparation techniques and implant macro-geometry on primary stability: An in vitro study. *Medicina Oral, Patologia Oral y Cirugia Bucal*. 2017; 22:e201-e06.
 40. Pocaterra A, Caruso S, Bernardi S, Scagnoli L, Continenza MA, Gatto R. Effectiveness of platelet-rich plasma as an adjunctive material to bone graft: a systematic review and meta-analysis of randomized controlled clinical trials. *International Journal of Oral and Maxillofacial Surgery*. 2016;45:1027-34.
 41. Giuca MR, Pasini M, Giuca G, Caruso S, Necozone S, Gatto R. Investigation of periodontal status in type 1 diabetic adolescents. *European journal of paediatric dentistry : official journal of European Academy of Paediatric Dentistry*. 2015;16:319-23.
 42. Giuca MR, Pasini M, Caruso S, Tecco S, Necozone S, Gatto R. Index of orthodontic treatment need in obese adolescents. *International Journal of Dentistry*. 2015, 2015.
 43. Caruso S, Sgolastra F, Gatto R. Dental pulp regeneration in paediatric dentistry: The role of stem cells. *European Journal of Paediatric Dentistry*. 2014;15:90-94.
 44. Marrelli M, Pujia A, Palmieri F, et al. Innovative approach for the in vitro research on biomedical scaffolds designed and customized with CAD-CAM technology. *International Journal of Immunopathology and Pharmacology*. 2016;29:778-83.
 45. Branemark PI, Hansson BO, Adell R, Breine U, Lindstrom J, Hallen O, Ohman A. Osseointegrated implants in the treatment of the edentulous jaw. Experience from a 10-year period. *Scand J Plast Reconst Surg Suppl*. 1977;16:1-132.
 46. Branemark PI. Osseointegration and its experimental background. *J Prosthet Dent*. 1983;50:399-410.
 47. Albrektsson T, Branemark PI, Hansson HA, Lindstrom J. Osseointegrated titanium implants. Requirements for ensuring a long-lasting, direct bone-to-implant anchorage in man. *Acta Orthop Scand*. 1981;52:155-70.
 48. Milillo L, Fiandaca C, Giannoulis F, Ottria L, Lucchese A, Silvestre F, Petruzzi M. Immediate vs non-immediate loading post-extractive implants: A comparative study of Implant Stability Quotient (ISQ). *Oral Implantol (Rome)*. 2016;9:123-31.
 49. Esposito M, Grusovin MG, Maghaireh H, Worthington HV. Interventions for replacing missing teeth: different times for loading dental implants. *Cochrane Database Syst Rev*. 2013;3:CD003878.
 50. Enriquez-Sacristan C, Barona-Dorado C, Calvo-Guirado JL, Leco-Berrocal I, Martinez-Gonzalez JM. Immediate post-extraction implants subject to immediate loading: a meta-analytic study. *Med Oral Patol Oral Cir Bucal*. 2011;16:e919-24.
 51. Sanz-Sanchez I, Sanz-Martin I, Figuero E, Sanz M. Clinical efficacy of immediate implant loading protocols compared to conventional loading depending on the type of the restoration: a systematic review. *Clin Oral Implants Res*. 2015;26:964-82.
 52. Zhu Y, Zheng X, Zeng G, Xu Y, Qu X, Zhu M, Lu E. Clinical efficacy of early loading versus conventional loading of dental implants. *Sci Rep*. 2015; 5:15995.
 53. Szmukler-Moncler S, Piattelli A, Favero GA, Dubruille JH. Considerations preliminary to the application of early and immediate loading protocols in dental implantology. *Clin Oral Implants Res*. 2000;11:12-25.
 54. Szmukler-Moncler S, Salama H, Reingewirtz Y, Dubruille JH. Timing of loading and effect of micromotion on bone-dental implant interface: review of experimental literature. *J Biomed Mater Res*. 1998;43: 192-203.
 55. Papaspyridakos P, Chen CJ, Chuang SK, Weber HP. Implant loading protocols for edentulous patients with fixed prostheses: a systematic review and meta-analysis. *Int J Oral Maxillofac Implants*. 2014; 29 Suppl:256-70.
 56. Javed F, Ahmed HB, Crespi R, Romanos GE. Role of primary stability for successful osseointegration of dental implants: Factors of influence and evaluation. *Interv Med Appl Sci*. 2013;5:162-7.
 57. Hui E, Chow J, Li D, Liu J, Wat P, Law H. Immediate provisional for single-tooth implant replacement with Branemark system: preliminary report. *Clin Implant Dent Relat Res*. 2001;3:79-86.
 58. Lorenzoni M, Pertl C, Zhang K, Wimmer G, Wegscheider WA. Immediate loading of single-tooth implants in the anterior maxilla. Preliminary results after one year. *Clin Oral Implants Res*. 2003; 14:180-7.
 59. Benic GI, Mir-Mari J, Hammerle CH. Loading protocols for single-implant crowns: a systematic review and meta-analysis. *Int J Oral Maxillofac Implants*. 2014;29 Suppl:222-38.

60. Pieri F, Aldini NN, Fini M, Corinaldesi G. Immediate occlusal loading of immediately placed implants supporting fixed restorations in completely edentulous arches: a 1-year prospective pilot study. *J Periodontol.* 2009;80:411-21.
61. Sul YT, Johansson CB, Jeong Y, Wennerberg A, Albrektsson T. Resonance frequency and removal torque analysis of implants with turned and anodized surface oxides. *Clin Oral Implants Res.* 2002;13:252-9.
62. Ertugrul AS, Tekin Y, Alpaslan NZ, Bozoglan A, Sahin H, Dikilitas A. Comparison of peri-implant crevicular fluid levels of adrenomedullin and human beta defensins 1 and 2 from mandibular implants with different implant stability quotient levels in nonsmoker patients. *J Periodontal Res.* 2014; 49:480-8.
63. Ramakrishna R, Nayar S. Clinical assessment of primary stability of endosseous implants placed in the incisor region, using resonance frequency analysis methodology: an in vivo study. *Indian J Dent Res.* 2007;18:168-72.
64. Hammerle CH, van Steenberghe D. The first EAO Consensus Conference 16-19 February 2006, Pfaffikon, Switzerland. *Clin Oral Implants Res.* 2006;17 Suppl 2:1.
65. Turkyilmaz I, Tozum TF, Tumer C, Ozbek EN. Assessment of correlation between computerized tomography values of the bone, and maximum torque and resonance frequency values at dental implant placement. *J Oral Rehabil.* 2006;33:881-8.
66. Misch CE. Density of bone: effect on treatment plans, surgical approach, healing, and progressive bone loading. *Int J Oral Implantol.* 1990;6:23-31.
67. Crespi R, Cappare P, Gherlone E, Romanos GE. Immediate occlusal loading of implants placed in fresh sockets after tooth extraction. *Int J Oral Maxillofac Implants.* 2007;22:955-62.
68. Cannizzaro G, Leone M, Esposito M. Immediate functional loading of implants placed with flapless surgery in the edentulous maxilla: 1-year follow-up of a single cohort study. *Int J Oral Maxillofac Implants.* 2007;22: 87-95.
69. Malo P, Rangert B, Nobre M. "All-on-Four" immediate-function concept with Branemark System implants for completely edentulous mandibles: a retrospective clinical study. *Clin Implant Dent Relat Res.* 2003;5 Suppl 1:2-9.
70. Tealdo T, Bevilacqua M, Menini M, Pera F, Ravera G, Drago C, Pera P. Immediate versus delayed loading of dental implants in edentulous maxillae: a 36-month prospective study. *Int J Prosthodont.* 2011;24:294-302.
71. Summers RB. A new concept in maxillary implant surgery: the osteotome technique. *Compendium.* 1994;15: 152, 54-6, 58 passim; quiz 62.
72. Crespi R, Cappare P, Gherlone E, Romanos GE. Immediate versus delayed loading of dental implants placed in fresh extraction sockets in the maxillary esthetic zone: a clinical comparative study. *Int J Oral Maxillofac Implants.* 2008;23:753-8.
73. Weber HP, Morton D, Gallucci GO, Rocuzzo M, Cordaro L, Grutter L. Consensus statements and recommended clinical procedures regarding loading protocols. *Int J Oral Maxillofac Implants.* 2009;24 Suppl: 180-3.
74. Spinelli D, Ottria L, De Vico GD, Bollero R, Bartattani Jr A, Bollero P. Full rehabilitation with nobel clinician® and provera implant bridge®: Case report. *ORAL and Implantology.* 2013;6:25-36.
75. O'Sullivan D, Sennerby L, Meredith N. Influence of implant taper on the primary and secondary stability of osseointegrated titanium implants. *Clin Oral Implants Res.* 2004;15:474-80.
76. Davies JE. Mechanisms of endosseous integration. *Int J Prosthodont.* 1998;11:391-401.
77. Degidi M, Nardi D, Piattelli A. Immediate restoration of small-diameter implants in cases of partial posterior edentulism: a 4-year case series. *J Periodontol.* 2009; 80:1006-12.
78. Penarrocha-Oltra D, Covani U, Penarrocha-Diago M, Penarrocha-Diago M. Immediate loading with fixed full-arch prostheses in the maxilla: review of the literature. *Med Oral Patol Oral Cir Bucal.* 2014;19:e512-7.
79. Yamada K, Hoshina H, Arashiyama T, et al. Immediate implant loading following computer-guided surgery. *J Prosthodont Res.* 2011;55:262-5.
80. Abad-Gallegos M, Gomez-Santos L, Sanchez-Garces MA, Pinera-Penalva M, Freixes-Gil J, Castro-Garcia A, Gay-Escoda C. Complications of guided surgery and immediate loading in oral implantology: a report of 12 cases. *Med Oral Patol Oral Cir Bucal.* 2011;16:e220-4.
81. Mundt T, Al Jaghsi A, Schwahn B, et al. Immediate versus delayed loading of strategic mini dental implants for the stabilization of partial removable dental prostheses: a patient cluster randomized, parallel-group 3-year trial. *BMC Oral Health.* 2016; 17:30.
82. Eliasson A, Blomqvist F, Wennerberg A, Johansson A. A retrospective analysis of early and delayed loading of full-arch mandibular prostheses using three different implant systems: clinical results with

- up to 5 years of loading. *Clin Implant Dent Relat Res.* 2009;11:134-48.
83. Fischer K, Stenberg T, Hedin M, Sennerby L. Five-year results from a randomized, controlled trial on early and delayed loading of implants supporting full-arch prosthesis in the edentulous maxilla. *Clin Oral Implants Res.* 2008;19:433-41.
 84. Lauritano D, Martinelli M, Mucchi D, Palmieri A, Muzio LL, Carinci F. Bacterial load of periodontal pathogens among Italian patients with chronic periodontitis: A comparative study of three different areas. *Journal of Biological Regulators and Homeostatic Agents.* 2016;30:149-54.
 85. Lauritano D, Scapoli L, Mucchi D, Cura F, Muzio LLO, Carinci F. Infectogenomics: Lack of association between *vdr*, *il6*, *il10* polymorphisms and “red Complex” bacterial load in a group of Italian adults with chronic periodontal disease. *Journal of Biological Regulators and Homeostatic Agents.* 2016;30:155-60.
 86. Checchi L, Gatto MR, Checchi V, Carinci F. Bacteria prevalence in a large Italian population sample: A clinical and microbiological study. *Journal of Biological Regulators and Homeostatic Agents.* 2016;30:199-208.
 87. Meynardi F, Pasqualini ME, Rossi F, Dal Carlo L, Biancotti P, Carinci F. Correlation between dysfunctional occlusion and periodontal bacterial profile. *J Biol Regul Homeost Agents.* 2016;30:115-21.
 88. Lombardo L, Carinci F, Martini M, Gemmati D, Nardone M, Siciliani G. Quantitative evaluation of dentin sialoprotein (DSP) using microbeads - A potential early marker of root resorption. *ORAL and Implantology.* 2016;9:132-42.
 89. Lauritano D, Cura F, Candotto V, Gaudio RM, Mucchi D, Carinci F. Evaluation of the Efficacy of Titanium Dioxide with Monovalent Silver Ions Covalently Linked (Tiab) as an Adjunct to Scaling and Root Planing in the Management of Chronic Periodontitis Using Pcr Analysis: A Microbiological Study. *J Biol Regul Homeost Agents.* 2015;29:127-30.
 90. Scapoli L, Girardi A, Palmieri A, Martinelli M, Cura F, Lauritano D, Carinci F. Quantitative Analysis of Periodontal Pathogens in Periodontitis and Gingivitis. *J Biol Regul Homeost Agents.* 2015;29:101-10.
 91. Lauritano D, Cura F, Candotto V, Gaudio RM, Mucchi D, Carinci F. Periodontal Pockets as a Reservoir of *Helicobacter Pylori* Causing Relapse of Gastric Ulcer: A Review of the Literature. *J Biol Regul Homeost Agents.* 2015;29:123-6.
 92. Scapoli L, Girardi A, Palmieri A, et al. Interleukin-6 Gene Polymorphism Modulates the Risk of Periodontal Diseases. *J Biol Regul Homeost Agents.* 2015;29:111-6.
 93. Carinci F, Girardi A, Palmieri A, et al. LAB®-Test 1: Peri-Implantitis and bacteriological analysis. *European Journal of Inflammation.* 2012;10:91-93.
 94. Carinci F, Girardi A, Palmieri A, et al. LAB®-test 2: Microflora and periodontal disease. *European Journal of Inflammation.* 2012;10:95-98.
 95. Carinci F, Girardi A, Palmieri A, et al. Lab®-test 3: Genetic susceptibility in periodontal disease. *European Journal of Inflammation.* 2012;10:99-101.
 96. Scapoli L, Girardi A, Palmieri A, et al. IL6 and IL10 are genetic susceptibility factors of periodontal disease. *Dent Res J (Isfahan).* 2012;9:S197-201.
 97. Carinci F, Girardi A, Palmieri A, et al. Lab-test 2: microflora and periodontal disease. *European Journal of Inflammation.* 2012;10:95-98.
 98. Cura F, Palmieri A, Girardi A, Martinelli M, Scapoli L, Carinci F. Lab-Test (R) 4: Dental caries and bacteriological analysis. *Dent Res J (Isfahan).* 2012;9:S139-41.
 99. Roncati M, Lauritano D, Cura F, Carinci F. Evaluation of light-emitting diode (led-835 nm) application over human gingival fibroblast: An in vitro study. *Journal of Biological Regulators and Homeostatic Agents.* 2016;30:161-67.
 100. Caccianiga G, Rey G, Paiusco A, Lauritano D, Cura F, Ormianer Z, Carinci F. Oxygen high level laser therapy is efficient in treatment of chronic periodontitis: A clinical and microbiological study using PCR analysis. *Journal of Biological Regulators and Homeostatic Agents.* 2016;30:87-97.
 101. Lauritano D, Bignozzi CA, Pazzi D, Palmieri A, Gaudio RM, Di Muzio M, Carinci F. Evaluation of the efficacy of a new oral gel as an adjunct to home oral hygiene in the management of chronic periodontitis. A microbiological study using PCR analysis. *J Biol Regul Homeost Agents.* 2016;30:123-8.
 102. Carinci F, Palmieri A, Girardi A, Cura F, Lauritano D. Aquolab ® ozone-therapy is an efficient adjuvant in the treatment of chronic periodontitis: A case-control study. *Journal of Orofacial Sciences.* 2015;7:27-32.
 103. Lauritano D, Cura F, Gaudio RM, Pezzetti F, Andreasi Bassi M, Carinci F. Polymerase Chain Reaction to Evaluate the Efficacy of Silica Dioxide Colloidal Solutions in the Treatment of Chronic Periodontitis: A Case Control Study. *J Biol Regul Homeost Agents.* 2015;29:131-5.

104. Lauritano D, Petruzzi M, Nardi GM, Carinci F, Minervini G, Di Stasio D, Lucchese A. Single Application of a Dessicating Agent in the Treatment of Recurrent Aphthous Stomatitis. *J Biol Regul Homeost Agents*. 2015;29:59-66.
105. Carinci F, Lauritano D, Cura F, Lopez MA, Bassi MA, Confalone L, Pezzetti F. Prevention of bacterial leakage at implant-Abutment connection level: An in vitro study of the efficacy of three different implant systems. *Journal of Biological Regulators and Homeostatic Agents*. 2016;30:69-73.
106. El Haddad E, Gianni AB, Mancini GE, Cura F, Carinci F. Implant-abutment leaking of replace conical connection nobel biocare® implant system. An in vitro study of the microbiological penetration from external environment to implant-abutment space. *ORAL and Implantology*. 2016;9:76-82.
107. Mancini GE, Gianni AB, Cura F, Ormanier Z, Carinci F. Efficacy of a new implant-abutment connection to minimize microbial contamination: An in vitro study. *ORAL and Implantology*. 2016;9:99-105.
108. Roncati M, Lucchese A, Carinci F. Non-Surgical treatment of peri-Implantitis with the adjunctive use of an 810-nm diode laser. *Journal of Indian Society of Periodontology*. 2013;17:812-15.
109. Scarano A, Tripodi D, Carinci F, Piccolomini R, D'Ercole S. Biofilm formation on titanium alloy and anatase-Bactercline® coated titanium healing screws: An in vivo human study. *Journal of Osseointegration*. 2013;5:8-12.
110. Brunelli G, Carinci F, Zollino I, Candotto V, Scarano A, Lauritano D. Sem evaluation of 10 infected implants retrieved from man. *European Journal of Inflammation*. 2012;10:7-12.
111. Scarano A, Sinjari B, Di Orio D, Murmura G, Carinci F, Lauritano D. Surface analysis of failed oral titanium implants after irradiated with ErCr:ysgg 2780 laser. *European Journal of Inflammation*. 2012;10:49-54.
112. Brunelli G, Carinci F, Zollino I, Candotto V, Scarano A, Lauritano D. Peri-implantitis. A case report and literature review. *European Journal of Inflammation*. 2012;10:1-5.
113. Scarano A, Piattelli A, Polimeni A, Di Iorio D, Carinci F. Bacterial adhesion on commercially pure titanium and anatase-coated titanium healing screws: An in vivo human study. *Journal of Periodontology*. 2010;81:1466-71.
114. Grecchi F, Zollino I, Candotto V, et al. A case of mandible osteonecrosis after a severe periimplant infection. *Dent Res J (Isfahan)*. 2012;9:S233-6.
115. Carinci F, Farina A, Zanetti U, et al. Alveolar ridge augmentation: a comparative longitudinal study between calvaria and iliac crest bone grafts. *J Oral Implantol*. 2005;31:39-45.
116. Carinci F, Pezzetti F, Volinia S, et al. Analysis of MG63 osteoblastic-cell response to a new nanoporous implant surface by means of a microarray technology. *Clinical Oral Implants Research*. 2004;15:180-86.
117. Oliveira DP, Palmieri A, Carinci F, Bolfarini C. Osteoblasts behavior on chemically treated commercially pure titanium surfaces. *J Biomed Mater Res A*. 2014;102:1816-22.
118. Andreasi Bassi M, Lopez MA, Confalone L, Carinci F. Hydraulic sinus lift technique in future site development: clinical and histomorphometric analysis of human biopsies. *Implant Dent*. 2015;24:117-24.
119. El Haddad E, Lauritano D, Carinci F. Interradicular septum as guide for pilot drill in postextractive implantology: a technical note. *J Contemp Dent Pract*. 2015;16:81-4.
120. Prasant MC, Thukral R, Kumar S, Sadrani SM, Baxi H, Shah A. Assessment of Various Risk Factors for Success of Delayed and Immediate Loaded Dental Implants: A Retrospective Analysis. *J Contemp Dent Pract*. 2016;17:853-56.
121. Schimmel M, Srinivasan M, Herrmann FR, Muller F. Loading protocols for implant-supported overdentures in the edentulous jaw: a systematic review and meta-analysis. *Int J Oral Maxillofac Implants*. 2014;29 Suppl:271-86.
122. De Vico G, Ottria L, Bollero P, et al. Aesthetic and functionality in fixed prosthodontic: sperimental and clinical analysis of the CAD-CAM systematic 3Shape. *Oral Implantol (Rome)*. 2008;1:104-115.
123. Moretto D, Gargari M, Nordsjo E, et al. Immediate loading: a new implant technique with immediate loading and aesthetics: Nobel Active. *Oral Implantol (Rome)*. 2008;1:50-55.
124. Spinelli D, De Vico G, Condò R, et al. Transcrestal guided sinus lift without grafting materials: A 36 months clinical prospective study. *ORAL Implantol (Rome)*. 2015;8:74-86.
125. Bartuli FN, Luciani F, Caddeo F, et al. Piezosurgery vs High Speed Rotary Handpiece: a comparison between the two techniques in the impacted third molar surgery. *Oral Implantol (Rome)*. 2013;6:5-10.
126. Baj A, Lo Muzio L, Lauritano D, et al. Success of immediate versus standard loaded implants: a short literature review. *Journal of biological regulators*

- and homeostatic agents. 2016;30(2 Suppl 1):183-8.
127. Lauritano D, Avantaggiato A, Candotto V, et al. Effect of somatostatin on dental pulp stem cells. *Journal of biological regulators and homeostatic agents*. 2015;29(3 Suppl 1):54-58.
128. Baj A, Sollazzo V, Lauritano D, et al. Lights and shadows of bone augmentation in severe resorbed mandible in combination with implant dentistry. *Journal of biological regulators and homeostatic agents*. 2016;30(2 Suppl 1):177-182.
129. Grecchi F, Perale G, Candotto V, et al. Reconstruction of the zygomatic bone with Smartbone®: case report. *Journal of biological regulators and homeostatic agents*. 2015;29(3 Suppl 1):42-47.
130. Lauritano D, Avantaggiato A, Candotto V, et al. Insulin activity on dental pulp stem cell differentiation: an in vitro study. *Journal of biological regulators and homeostatic agents*. 2015;29(3 Suppl 1):48-53.
131. Baj A, Trapella G, Lauritano D, et al. An overview on bone reconstruction of atrophic maxilla: success parameters and critical issues. *Journal of biological regulators and homeostatic agents*. 2016;30(2 Suppl 1):209-215.
132. Tettamanti L, Bassi MA, Trapella G, et al. Applications of biomaterials for bone augmentation of jaws: clinical outcomes and in vitro studies. *Oral Implantol (Rome)*. 2017 Apr 10;10(1):37-44. eCollection 2017 Jan-Mar.

Correspondence to:

Lucia Tettamanti
Department of Medicine and Surgery
University of Insubria
Via Piatti 10
21100 Varese, Italy
Phone: +39.0332-825625; Fax: +39.0332-825655
E-mail: lucia.tettamanti@uninsubria.it