# Immediate Placement or Immediate Restoration/Loading of Single Implants for Molar Tooth Replacement: A Systematic Review and Meta-analysis

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Purpose: This systematic review and meta-analysis assessed the survival of immediately placed single implants in fresh molar extraction sites and immediately restored/loaded single molar implants in healed molar sites. Materials and Methods: A search of the main electronic databases, including the Cochrane Oral Health Group's Trials Register, was conducted up to November 1, 2008. The meta-analysis was prepared in accordance with the guidelines of the Academy of Osseointegration Workshop on the State of the Science on Implant Dentistry. The data were analyzed with statistical software. Results: For immediately placed molar implants, nine studies describing 1,013 implants were included to support a survival rate of 99.0%. There were no significant differences between immediate and delayed loading/restoration in molar sites (relative risk of 0.30, 95% confidence interval 0.05 to 1.61; P = .16). For immediate restoration/loading of single implants in healed molar sites, seven studies with 188 single implants were identified. In this case, the implant survival rate was 97.9%, with no difference between immediate and delayed loading (relative risk of 3.0, 95% confidence interval: 0.33 to 27.16; P = .33). Favorable marginal bone level changes in the immediate loading group were detected at 12 months (mean difference of -0.31, 95% confidence interval: -0.53 to -0.096; P = .005). Conclusions: The protocols of immediate placement and immediate restoration/loading of single implants in mandibular molar regions showed encouraging results. Int J Oral Maxillofac Implants 2010;25:401-415

**Key words:** dental implants, immediate loading, immediate placement, immediate restoration, meta-analysis, single implant, systematic review

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In view of the original conventional approach to the surgical placement of oral implants in edentulous patients, clinicians historically allowed extraction sockets to heal prior to implant placement. Following this, a second procedure was required to place the implant transgingivally<sup>2</sup> or subgingivally<sup>3</sup>; in the latter case, a third surgical appointment was necessary to expose the implant for restoration. These protocols were less technically demanding but subjected patients to multiple surgical interventions. Furthermore, implant loading, regardless of the type of prosthesis, was traditionally delayed until after a conventional healing period to allow for osseointegration. Recently, these traditional protocols for placing and loading oral implants, especially in cases of single tooth replacement, have been revised to meet subjective and objective requirements for fewer surgical interventions and shorter implant treatment times.

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| Table 1 Definitions   | of Placement and Loading Protocols <sup>4,19</sup>  |
|-----------------------|---|
| Protocol              | Definition  |
| Placement             |   |
| Immediate placement   | Implants placed in fresh extraction sockets (type I)4   |
| Delayed placement     | Implants placed in healed sites after at least 4 months of healing (type IV) <sup>4</sup>                     |
| Loading               |   |
| Immediate restoration | Restoration placed within 48 hours of implant placement but not in centric or eccentric occlusal contact with |
|                       | the opposing dentition during healing   |
| Immediate loading     | Restoration placed into occlusal load within 48 hours after implant placement                                 |
| Conventional loading  | Restoration placed in a second procedure after a healing period of 3 to 6 months                              |

The term immediate implant placement refers to the placement of oral implants at the time of tooth extraction.4 This protocol offers several advantages, including a reduction in the number of surgical procedures,5,6 preservation of esthetics and bone height and width, 6,7 improved quality of life, and increased patient comfort and satisfaction.8 Moreover, the individual healing potential of the fresh extraction socket as well as the implant surface characteristics may provide better opportunities for osseointegration.5,7,9 With careful patient selection, immediate implant placement, particularly in the esthetic zone of the mouth, has gained acceptance among the scientific community, with several authors showing that success rates can be achieved that are similar to those obtained for delayed implants placed into healed extraction sites. 10-12

Therefore, immediate implant placement is currently a very popular choice to replace a missing single tooth in the esthetic zone of the mouth. 13 However, the immediate placement of a single implant in molar regions involves numerous challenges related to site-specific anatomic, occlusal, and biomechanical factors. The reported long-term success of oral implants placed in healed maxillary and mandibular molar regions is inferior to that of implants placed in anterior esthetic sites. 14,15 The possibility of predictable outcomes with immediate implantation in molar sites is additionally compromised because of the larger extraction sockets, poor quality of bone (particularly in the maxillary molar regions<sup>16,17</sup>), and less bone apical to the socket because of the proximity of the maxillary sinus or inferior dental canal.18

In addition to immediate placement, single implants may also be restored/loaded with an implant-supported prosthesis immediately after placement, with the objectives of shortening treatment time, improving esthetics, and meeting patients' expectations. Immediate loading is defined as placement of a restoration in occlusion within the first 48 hours following surgical implant placement.<sup>19</sup>

Furthermore, other authors distinguish between immediate nonfunctional restoration and immediate functional loading, depending on whether a restoration is placed into occlusion (Table 1).<sup>4,19,20</sup>

Several specific studies have therefore reported high survival rates with immediate restoration/loading or early restoration/loading of single implants in the anterior region, albeit in the short term.<sup>21–25</sup> With recent innovations in implant designs and surface characteristics, immediate loading of oral implants can also be achieved in the posterior regions for short-term periods.<sup>26–30</sup> Several extensive reviews have demonstrated both the effectiveness and the limitations of immediate placement or immediate restoration/loading of oral implants.<sup>31–44</sup>

Although there is a trend to combine the two protocols into immediate placement with immediate loading, the current literature is still reporting either immediate placement or immediate restoration/loading in the molar region. Therefore, it was deemed important to systematically review the placement of single implants in fresh molar extraction sockets and the immediate restoration/loading of single implants placed in healed molar sites in a single paper. The aims of this systematic review were twofold: first, to evaluate the overall effectiveness of single implant placement in fresh molar extraction sockets and, second, to summarize the clinical outcomes of immediate restoration/loading of single implants in healed molar sites.

### **MATERIALS AND METHODS**

This systematic review was conducted according to the guidelines of the Academy of Osseointegration (AO) Workshop on the State of the Science in Implant Dentistry (SSID).<sup>45,46</sup> A PICO format<sup>47</sup> was constructed to identify the objectives and the inclusion criteria. The essential four elements (**P**opulation, **I**ntervention, **C**omparison, **O**utcome) were summarized in the following two questions:

- Participant: Patients who needed implant placement immediately following extraction of a single molar tooth. Intervention: Immediate implant placement in molar extraction sockets. Comparison: Delayed implant placement in healed molar ridges. Outcome: Implant failure rate.
- Participant: Patients that needed immediate restoration/loading of a single implant in a healed molar region. Intervention: Immediate implant restoration/loading in healed molar ridges. Comparison: Delayed implant restoration/loading in healed molar ridges. Outcome: Implant failure rate.

Early placement and loading were not included in this review; because the current definition is not clear, it is difficult to compare the outcomes of studies available in the literature. 4,19

### Search Methodology

A comprehensive literature search of the following databases was conducted:

- MEDLINE (1966 through October 2008)
- EMBASE (1980 through October 2008)
- The Cochrane Oral Health Group's Trials Register (up to October 2008)
- The Cochrane Central Register of Controlled Trials (CENTRAL)
- · United Kingdom National Research Register
- Australian New Zealand Clinical Trials Registry
- ISI Proceedings for relevant conference abstracts

The search involved human studies reported in any language, with two search strategies being performed. For the first PICO question (regarding the immediate placement of single implants in molar extraction sites), the following combinations of search terms and key words were used: ("immediate placement" OR "immediate implant" OR "immediate implantation") AND ("single implant" OR "single tooth implant" OR "single oral implant") AND ("molar extraction socket" OR molar extraction site") AND ("survival rate" OR "success rate") AND ("randomized controlled clinical trials" OR "controlled clinical trials" OR · "prospective" OR "cohort studies" OR "longitudinal" OR "retrospective"). For the second PICO question (regarding the immediate restoration/loading of single implants in healed molar sites), the following combinations of search terms and key words were used: ("immediate loading" OR "immediate restoration") AND ("single implant" OR "single tooth implant" OR "single oral implant") AND ("molar sites" OR "molar regions") AND ("survival rate" OR "success rate") AND ("randomized controlled clinical trials" OR "controlled clinical trials" OR "prospective" OR "cohort studies" OR "longitudinal" OR "retrospective").

Furthermore, bibliographies of the selected articles and relevant reviews were manually searched. A hand search was also conducted in the following journals for the past 5 years (up to November 2008): Clinical Implant Dentistry and Related Research, Clinical Oral Implants Research, European Journal of Oral Implantology, Implant Dentistry, International Journal of Oral & Maxillofacial Implants, International Journal of Periodontics & Restorative Dentistry, International Journal of Prosthodontics, Journal of Clinical Periodontology, Journal of Dental Research, Journal of Esthetic and Restorative Dentistry, Journal of Oral Implantology. Journal of Oral Rehabilitation, Journal of Periodontal Research, Journal of Periodontology, Journal of Prosthetic Dentistry, Journal of Prosthodontics, Oral Surgery Oral Medicine Oral Pathology Oral Radiology and Endodontics, and Quintessence International. In addition, the reviewers attempted to contact corresponding authors, where appropriate, to confirm data extraction and/or obtain missing data.

### Study Selection

The searches were carried out by two authors (MA and AP) independently. Any disagreements were resolved by discussion and, if necessary, by a third author (WD). All types of study designs were included, except for case reports because of their lack of quantitative outcomes.<sup>44</sup> In addition, the following inclusion criteria for study selection were considered:

- A minimum of 10 single implants immediately placed or restored/loaded in the maxillary or mandibular molar region
- A mean follow-up period of at least 6 months
- A clearly presented survival or success rate, or available data that allowed calculation of survival or success rate
- The use of endosseous solid root-form titanium oral implants

Implant success and survival were defined according to the broad criteria proposed at the AO workshop to include more studies in the review.<sup>45</sup> Classifications from previous consensus meetings<sup>4,19</sup> were adopted to define placement and loading protocols.

### **Data Abstraction**

The following information was retrieved from the selected studies using a specially designed data template:

 Publication details (title, author(s), journal, year, volume, issue number, pages)

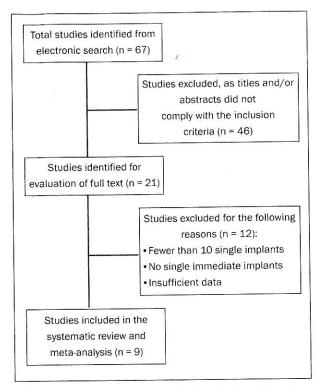


Fig 1 Search strategy for studies related to the immediate placement of single implants in molar extraction sockets.

- · Type of study
- Patient details
- Number of implants immediately placed in maxillary or mandibular molar extraction sockets
- Number of implants immediately restored/loaded
- · Details of the surgical approach
- Type of bone grafting material, if used
- Implant survival or success rate of treatment group(s)
- · Follow-up period

### **Quality Assessment**

Six quality categories<sup>46</sup> were used to evaluate the quality of each selected study according to its design: "fair" for a retrospective study, "average" for a prospective case study, "good" for a prospective study with historical controls, "better" for a prospective study with concurrent controls, "best" for a double-blind randomized controlled trial (RCT), and "unknown" when the study design could not be ascertained or fit none of the definitions.

### **Statistical Methods**

The pooled proportion of implant survival or success was estimated along with the 95% confidence intervals (CIs) using R Statistical Software (version 2.7.1, R

Foundation for Statistical Computing). The random-effect pooling model of DerSimonian and Laird<sup>48</sup> was employed for comparison of heterogenous studies. For controlled studies, relative risk and 95% Cls were used to calculate the effect of immediate placement of single implants in fresh molar extraction sockets compared to placement in healed ridges. When heterogeneity was not significant, the Mantel-Haenszel method<sup>49</sup> was used to calculate the fixed-effects pooled relative risks.

The impact of between-study heterogeneity was evaluated using the chi-square test. Because this method has low sensitivity, a P value of < .1 was considered indicative of significant heterogeneity.<sup>50</sup> The variation across studies because of heterogeneity was assessed using the  $I^2$  statistic,<sup>51</sup> where an  $I^2$  value of > 50% was considered significant for the presence of heterogeneity.

### **RESULTS**

## **Immediate Placement in Extraction Sockets**

The initial electronic search identified 67 citations for reviewing. Of these, 46 were rejected after the abstracts were screened. Following the assessment of the full texts of the remaining 21 articles, 12 studies were excluded for the following reasons (Fig 1):

- Three studies were case series with fewer than 10 single implants.<sup>52–54</sup>
- Three studies failed to specify the total number of immediate single implants in molar sites. 55–57
- Two studies included both single and splinted immediate implants in molar sites; however, the number of single implants was less than 10.<sup>58,59</sup>
- One study failed to report the number of failed single implants in the immediate placement group.<sup>60</sup>
- One study included only two single implants in molar extraction sockets.<sup>61</sup>
- One study compared three different sinus augmentation procedures in conjunction with placement of single implants placed in healed sites.<sup>62</sup>
- One study excluded single implants inserted in molar extraction sockets.<sup>63</sup>

Manual searching did not provide any additional studies. A total of nine studies<sup>64–72</sup> including 1,013 immediate single implants in either maxillary and mandibular molar extraction sockets was included. Contact with the corresponding authors of two studies<sup>71,72</sup> provided additional relevant data and confirmed eligibility for inclusion. The characteristics of the included studies are summarized in Table 2.

| Table 2 Cha                                      | Characteristics of the Selected Studies or  | e Selected Stud                            | ies on Immedia              | te Placement of                                     | Immediate Placement of Single Implants in Molar Extraction Sockets | in Molar Extract         | ion Sockets              |   |                                 |
|--|---|--|-----------------------------|---|--|--------------------------|--------------------------|---|---------------------------------|
| Characteristic                                   | Becker and<br>Becker <sup>64</sup>          | Bianchi and<br>Sanfilippo 65               | Cafiero et al <sup>66</sup> | Fugazzotto <sup>67</sup>                            | Fugazzotto <sup>68</sup>   | Fugazzotto <sup>69</sup> | Fugazzotto <sup>70</sup> | Levin et al <sup>71</sup>                 | Peñarrocha et al <sup>72</sup>  |
| Study design<br>No. of participants/<br>implants | Retrospective<br>/ 22/24; IP: 13,<br>DP: 11 | RCT<br>116/42                              | Prospective<br>82/82        | Retrospective<br>319/33                             | Prospective<br>83/83   | Retrospective<br>386/387 | Prospective<br>320/341   | Retrospective<br>81/81; IP: 21,<br>DP: 60 | Retrospective<br>100/55;IP: 11, |
| Participant age                                  | 48 to 73                                    | 19 to 73                                   | 21 to 85                    | 24 to 86  | 38 to 68   | 28 to 76                 | 26 to 81                 | 18 to 74                                  | 20 to 76                        |
| Participants'                                    | Good health; bruxers                        | Absence of                                 | Medically fit,              | Absence of systemic                                 | No systemic con-   | No systemic con-         | No systemic con-         | The presence of                           | Healthy patients with           |
| inclusion criteria                               | and clenchers were                          | systematic problems,                       | , sufficient residual       | conditions, uncon-                                  | ditions, chemo-  | ditions, chemo-          | ditions, chemo-          | adjacent teeth,                           | sufficient alveolar             |
|  | excluded                                    | good oral hygiene,                         | alveolar bone               | trolled periodontal                                 | therapy, radiation   | therapy, radiation,      | therapy, radiation or    | and follow-up data                        | ridge height and                |
|  |   | stable occlusion,                          | volume, ≥; 2 mm             | disease, psycho-                                    | or smoking > 10  | or smoking > 10          | smoking > 10             | of 6 mo or more                           | width; smokers were             |
|  |   | < 15 cigarettes/day,<br>no alcohol or drug | of KT, full-mouth           | logical problems,                                   | cigarettes/d   | cigarettes/d             | cigarettes/d             |   | not excluded                    |
|  |   | dependency                                 | scores ≤ 25% at             |   |  |                          |                          |   |                                 |
|  |   |  | baseline, < 10              | 12-mo period  |  |                          |                          |   |                                 |
|  |   |  | cigarettes/d                |   |  |                          |                          |   |                                 |
| Implant system                                   | Brånemark System                            | Solid-screw ITI                            | TE ITI implants             | IMZ TPS cylindric                                   | TE ITI implants with   | TE ITI implants with     | TE ITI implants with     | Screw-type Core-                          | Defcon Avantblast-              |
|  | (machined-surface)                          | implants                                   | with SLA surface            | implants  | SLA surface  | SLA surface              | SLA surface              | Vent implants                             | surface implants                |
|  | (Nobelpharma)                               | (Straumann)                                | (Straumann)                 | (Biomet/Interpore),<br>TPS threaded<br>ITI implants | (Straumann)  | (Straumann)              | (Straumann)              | (Zimmer Dental)                           | (Impladent)                     |
| Implant diameter                                 | 3.75, 4.0, 5.0                              | 4.1  | 4.8                         | (Straumann)<br>3.3, 4.0, 4.25                       | 4.1  | 4.1, 4.8                 | 4.1, 4.8                 | s 4.0 (n = 24),                           | 5.5                             |
| (mm)   |   |  |                             |   |  |                          |                          | > 4.0 (n = 57)                            |                                 |
| Implant length<br>(mm)                           | 6, 8, 10, 13, 15                            | 10, 12, 14                                 | 8, 10, 12, 14               | 8, 11, 13, 15                                       | 11.8   | 8, 10, 12                | 8, 10, 12                | < 13 (n = 9),<br>$\ge 13 (n = 72)$        | 8.5, 10, 11.5, 13               |
| Implant location                                 | Maxilla, mandible                           | Maxilla, mandible                          | Maxilla, mandible           | Maxilla, mandible                                   | Maxilla  | Maxilla                  | Mandible                 | Mandible, maxilla                         | Mandible, maxilla               |
| Surgical protocol                                | Mesial-distal luxation                      | Test group: single                         | Full-thickness flap         | Atraumatic extraction,                              | Vertical and hori-   | Different flap designs,  | Sulcular incisions       | Atraumatic                                | Atraumatic                      |
|  | followed by gentle                          | immediate implants                         | ± releasing incisions,      |   | zontal releasing inci-   | sectioning of all        | with vertical and        | extraction, socket                        | extraction, preserving          |
|  | removal; implants                           | with connective                            | site preparation            | residual socket filled                              | sions, distal wedge  | maxillary molars, re-    | horizontal releasing     | debrided, sockets                         | the alveolar bone               |
|  | were placed either                          | tissue graft (n = 96);                     | in the area of inter-       | with DFDBA, area                                    | incisions for second   | moval of each root       | incisions, hemi-         | prepared with                             | and interdental                 |
| ,  | into one of the root                        | control group: single                      | radicular bone,             | covered with  | molars followed by   | individually; care to    | section/trisection of    | standard drills in                        | papillae                        |
|  | sockets or into the                         | immediate implants                         |                             | e-PTFE membrane                                     | sequential use of  | preserve interradic-     | mandibular molars,       | interdental bone                          |                                 |
|  | interradicular bone                         | without grafts (n = 20)                    | in the                      |   | osteotome to spread  | ular bone with           | socket debridement       | with maximum use                          |                                 |
|  |   |  | apical portion of           |   | the interradicular   | concomitant regen-       |                          | of bone apical to                         |                                 |
|  |   |  | interradicular septum       |   | bone and lift the sinus  | eration if necessary;    |                          | the extraction                            |                                 |
|  |   |  |                             |   |  | tapered osteotomes       |                          | sockets                                   |                                 |
|  |   |  |                             |   |  | were used to prepare     |                          |   |                                 |
|  |   |  |                             |   |  | the interradicular bone  |                          |   |                                 |

| Characteristic   |  |   |  |  |  |  |  |   |   |
|--|--|---|--|--|--|--|--|---|---|
|  | Becker and<br>Becker <sup>64</sup>   | Sanfilippo 65 (   | Caflero et al <sup>66</sup>  | Fugazzotto 67  | Fugazzotto <sup>68</sup>   | Fugazzotto <sup>69</sup>   | Fugazzotto <sup>70</sup>   | Levin et al <sup>71</sup>   | Peñarrocha et al <sup>72</sup>                              |
| Use of regenerative Not clear material   | Not clear  | Autologous connective Deproteinized tissue graft bovine bone mi particles (Bio-Calestile) + bio bable collagen brane (Bio-Gid Geistlich) (gaps Calestile) (gaps 1.0 mm) | neral<br>Jss,<br>resor-<br>mem-<br>e,<br>s, >  | Resorbable TCP (Miter) or DFDBA (Musculoskeletal Foundation) beneath e-PTFE membrane (W.L. Gore) | DFDBA (Exatech) and/<br>or osseous coagulum<br>and bioabsorbable or<br>titanium-reinforced<br>e-PTFE membrane<br>(W.L. Gore) secured<br>with two buccal<br>titanium tacks (Ace<br>Surgical Supply) | Bracked with bovine grafted with bowne grafted with bovine grafted with bowne bone matrix (Osteo-bone matrix (Osteo-health) or deminer-alized to impregnated with putty impregnated cortical chips (Exatech); Itlanium (Exatech) (n = 166 reinforced or biomembrane (WL. Gore) absorbable coverwas placed over membrane (WL. Gore) absorbable cover dareas (n = 371) was placed over dareas (n = 63) | HDD > 3 mm were HDD > 3 mm were I grafted with bovine grafted with bovine bone matrix (Osteo- bone matrix (Osteo- health) or deminer- health) (n = 165) or demineralized bone impregnated with putty impregnated cortical chips (Exatech); titanium. (Exatech) (n = 167); reinforced or bio- titanium-reinforced absorbable covering (n = 269) or bio- membrane (W.L. Gore) absorbable covering was placed over grafted areas (n = 371) was placed over grafted areas (n = 63)                         | Deproteinized bovine bone boune bone mineral particles (Bio-Oss, Geistlich) (reported in only one case) | Dehiscences or HDD > 2 mm were grafted with autologous bone |
| Use of antibiotics Time to loading Marginal bone level changes Observation period Implant survival | 2 g penicillin or erythromycin 2 h prior to surgery and 1 g daily for 7 d postop 4-6 mo Not measured 24 mo IP: 100, DP: 90.9 | Not clear 3-4 mo 0.75 mm for all groups at 12 mo 72 to 108 mo   | Systemic antibiotics (eg. amoxicillin) for 5 d postsurgically 3 mo -3.2 mm at 12 mo 12 | Postoperative antibiotic coverage Not clear Not measured 78 to 133 mo 93.9*                      | 500 mg amoxicillin<br>3× daily for 10 d<br>6 mo<br>Not measured<br>18 mo<br>100†   | 500 mg amoxicillin 500 mg am 3× daily for 10 d 3× daily for 10 d 3× daily for 10 d 33 mg 3× for amoxicillin- daily for 10 sensitive subjects) for amoxicillin- sensitive subjects) for amoxicillin- sensitive su 6 m 3-7 mo Not measured Not measured Not measure 40.3 mo 99.5† 99.5†  | 500 mg amoxicillin 500 mg amoxicillin Not clear 3× daily for 10 d 3× daily for reperter terythromycin, 333 mg 10 d (erythromycin, 333 mg 10 d (erythromycin, 31 mg 3× for amoxicillin-daily for 10 d, sensitive subjects) for amoxicillin-sensitive subjects) 3-7 mo 3-7 mo Not measured Not clear Not measured Not clear Pto.3 mo 6 to 125 in 10.0. E | Not clear<br>3-6 mo<br>Not clear<br>6 to 125 mo<br>IP: 100, DP: 90                                      | Not clear Not clear IP: 0.83 mm; DP: 0.85 mm 12 mo          |

DEDBA = demineralized freeze-dried bone allograft; DP = delayed placement; e-PTE = expanded polytetrafluoroethylene; HDD = horizontal defect dimension; IP = immediate placement; KT = keratinized tissue; RCT = randomized controlled trial; SLA = sandblasted, large-grit, and acid-etched; TCP = tricalcium phosphate; TE = tapered effect; TPS = titanium plasma-spray

\*Success rate (Albrektsson et al<sup>72</sup>); \*success rate (Albrektsson et al<sup>72</sup>); \*success rate (Buser et al<sup>75</sup>).

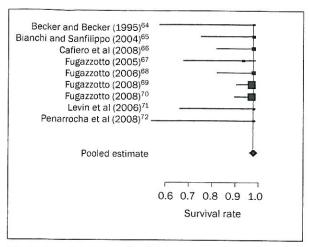


Fig 2 Forest plot analysis of the survival of single implants placed in molar extraction sockets.

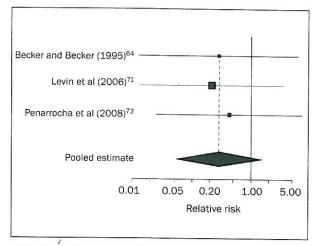


Fig 3 Forest plot analysis of the survival of single implants following immediate placement versus delayed placement in molar sites.

Description of the Included Studies. The main indications for extraction included periodontally or restoratively hopeless molars, root fracture, and endodontic failures. 64,68 All the selected studies excluded patients who had a bruxing/parafunctional habit, systemic diseases, or poor oral hygiene. Heavy smokers (> 10 cigarettes per day) were excluded from all studies but one.<sup>72</sup> In two studies,<sup>66,67</sup> molars with untreated periodontitis were also excluded from the study. Four studies<sup>66,68-70</sup> included in this review used tapered implants with sandblasted and acid-etched surfaces and a shoulder diameter of 6.5 mm. Moreover, most of the studies<sup>66-70,72</sup> included bone augmentation procedures to fill peri-implant voids. The use of preoperative and/or postoperative antibiotics was clearly described by six studies. 64,66-70 The interval before the implants were loaded ranged from 3 months<sup>66</sup> to 7 months.<sup>70</sup>

**Quality of the Included Studies.** Only one of the included studies was an RCT.<sup>65</sup> However, in this study the randomization was related to the use of connective tissue grafts rather than the time of implant placement. Three studies were categorized as average, as they were prospective case studies without historical or concurrent controls.<sup>66,68,70</sup> The remaining studies were retrospective and were classified as fair.<sup>64,67,69,71,72</sup>

Implant Success/Survival or Failure. All the studies reported the number of failures for implants placed immediately into molar extraction sockets. Three studies<sup>67-69</sup> used criteria published by Albrektsson et al<sup>73,74</sup> to determine the implant success rate, whereas one study<sup>72</sup> defined implant success according to the criteria of Buser et al.<sup>75</sup> The implant survival

rates ranged from 93.9%<sup>67</sup> to  $100\%^{64-66,68,71,72}$  over a period of 12 to 133 months, with an overall pooled estimate of 0.99 (random-effects model, 95% CI: 0.982 to 0.995; Fig 2) in both maxillary and mandibular molar sites. A high degree of statistical homogeneity between studies was detected (P = 1.0 with  $I^2 = 0\%$ ).

Only three studies<sup>64,71,72</sup> included separate failure rates for immediately placed single implants and those placed in healed sites; comparable results were reported for the two groups. Similarly, the meta-analysis did not reveal any significant difference between immediate and delayed placement of single implants in molar sites (relative risk of 0.30, 95% Cl: 0.05 to 1.61, P = .16; Fig 3). No significant statistical heterogeneity was observed between studies (P = 1.0 with  $I^2 = 0\%$ ).

# Immediate Restoration/Loading in Healed Molar Sites

The electronic search identified 129 articles for further consideration (Fig 4). After the abstracts and key words were screened, 21 studies were regarded as potentially eligible and retrieved for full-text analysis. Fourteen studies were excluded for the following reasons:

- Eight studies did not specify the number of single implants in the molar sites.<sup>27,31,76–81</sup>
- Three studies had molar implants restored with fixed partial dentures.<sup>82–84</sup>
- Two studies reported on early loading in posterior mandibular<sup>85</sup> and maxillary<sup>86</sup> sites.
- One study mentioned molar sites but involved only immediate loading of single implants in premolar areas.<sup>87</sup>

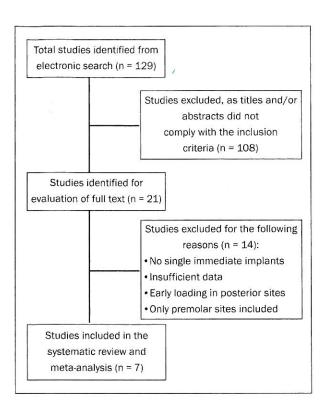


Fig 4 Search strategy for studies related to immediate restoration/loading of single implants in healed molar sites.

No additional relevant studies were identified from the hand search of journals and references. Overall, seven studies<sup>27–29,88–91</sup> with 188 implants in molar sites that had been immediately restored/loaded with single crowns met the inclusion criteria and were included in the analysis (Table 3).

Description of the Included Studies. All the studies<sup>27–29,88–91</sup> applied strict inclusion criteria for selecting participants, which included good general systemic health and adequate bone quality and quantity; smokers, bruxers, and patients with poor oral hygiene were excluded. Sites that required bone augmentation were excluded in two studies.<sup>29,90</sup> Five studies<sup>27,29,89–91</sup> reported immediate loading of single implants with crowns placed in molar sites, while only two<sup>28,88</sup> described the immediate restoration of single implants, also with crowns, in molar sites. Four studies<sup>27,29,89,90</sup> reported on provisional restoration with crowns in occlusion. In one study,91 lateral excursions were eliminated and the implants were restored in centric relation, while minimal contact in occlusion was maintained in another.88 Only one study28 reported an absence of full occlusal contact with the molar crowns in place, hence immediate restoration.

Only one study reported that single implants were placed in healed sites of both maxillary and mandibular molar sites.<sup>88</sup> The remaining studies<sup>27–29,89–91</sup> reported on single implants placed in healed mandibular molar ridges.

Quality of the Included Studies. Two studies<sup>29,91</sup> compared two different loading protocols, namely, immediate versus delayed loading. These studies were identified as RCTs and therefore categorized as better. The remaining studies<sup>27,28,88–90</sup> were prospective case studies without historical or concurrent controls and were classified as average. It is worth noting that the use of a blinded investigator to measure radiographic bone level was featured in only one study.<sup>91</sup>

Implant Success/Survival or Failure with Marginal Bone Level Changes. All the studies reported a high survival rate for immediately restored/loaded single implants in healed molar sites, ranging from  $90.9\%^{88}$  to  $100\%^{27,28,89}$  over an observation time of 6 to 36 months. The meta-analysis showed an overall pooled estimate of 0.979 (random-effects model, 95% CI: 0.947 to 0.991; Fig 5). No evidence of statistical heterogeneity was observed (P = 1.0) with  $I^2 = 0\%$ ).

Only two studies<sup>29,91</sup> compared implant survival and marginal bone level changes between two different loading protocols, namely immediate and conventional loading. In both studies, single implants were placed in mandibular molar sites and followed up for 12 months; both studies were described as RCTs. One implant failed in the immediate loading group in each study. In one study,<sup>29</sup> no significant differences in marginal bone level changes were observed between the

| Table 3 Characteristics of the Selected Studies  | stics of the Selec   |  | nediate Restoration  | n and/or Loading of  | Single Implants Pla   | sced in Healed Mola   | of Immediate Restoration and/or Loading of Single Implants Placed in Healed Molar Extraction Sockets  |
|--|--|--|--|--|---|---|---|
| Characteristic   | Rao and Benzi <sup>27</sup>  | Payer et al <sup>28</sup>  | Güncü et al <sup>29</sup>  | Abboud et al <sup>88</sup>   | Calandriello et al <sup>89</sup>  | Cornelini et al <sup>90</sup>   | Schincaglia et al <sup>91</sup>   |
| Study design No. of participants/implants Participant age range (y) Participants' inclusion criteria | THE PROPERTY OF THE PROPERTY OF THE PARTY OF | Prospective 24/19 29-63 Good general health, adequate bone width and height; exclusion criteria: bruxers, smokers, questionable patient cooperation  | IL: 12/12, CL: 12/12 30-55 Medically fit patients with bilateral loss of mandibular first molar, exclusion criteria: parafunctional habits, smoking, poor oral hygiene and sites where bone augmentation is needed | Prospective 20/11 NS Good general systemic health; exclusion criteria: daily smokers, bruxers, and those with uncontrolled periodontitis | Prospective 44/50 26-73 Absence of systematic problems, sufficient vertical bone height; bruxers were excluded, but smokers were not excluded | Prospective 30/30 27-59 Medically fit, sufficient I bone quality, exclusion criteria: parafunctional habits, poor oral hygiene, and sites where bone augmentation is needed | RCT IL: 15/15, CL: 15/15 35-68 Good general health, adequate bone width and height, no need for bone graft procedures, adequate implant stability, insertion torque ≥ 20 Ncm; smokers were not excluded |
| Implant system   | Replace Select TiUnite XIVE screw-type (threaded with oxidized implants (Dentsply surface, tapered body Friadent) and 1.5 mm-high collar) (Nobel Biocare)  | XiVE screw-type<br>implants (Dentsply/<br>Friadent)  | TiUnite MK IV implants,<br>anodized surface<br>(Nobel Biocare)   | Ankylos implants (sand-<br>blasted surface with 2-mm<br>smooth transmucosal<br>collar) (Friadent)  | TrUnite MK III Wide Platform implants (Nobel Biocare)   | ITI solid implants with sandblasted, acid-etched surface (Straumann)  | TiUnite MK III Wirde-Platform implants (Nobel Biocare)  |
| Implant diameter (mm)  | 4.3, 5.0, 6.0  | 5  | 4.0  | 3.5, 5.5   | 5.0   | 4.1, 4.8  | 5.0   |
| Implant length (mm)  | 10, 13   | 11, 13, 15   | 11.5   | 9.5, 11  | At least 10   | 10, 12  | 8.5, 10, 11.5   |
| Implant location   | Mandible   | Mandible   | Mandible   | Maxilla, mandible  | Mandible  | Mandible  | Mandible  |
| Type of occlusion  | L  | ₩.   | 1  | R  | =   | 7   | =   |
| Marginal bone level<br>changes (mm)  | 1.12 ± 1.06<br>(at 12 mo)  | 0.93 (at 6 mo)   | L: $0.45 \pm 0.39$ (at 12 mo); 0.01 (at 12 mo) ICL: $0.68 \pm 0.3$ (at 12 mo)  | 0.01 (at 12 mo)  | 1.3 ± 0.6 (at 12 mo)  | 0.22 (at 6 mo)  | IL: 0.77 ± 0.38 (at 12 mo),<br>CL: 1.2 ± 0.55 (at 12 mo)  |
| Implant stability measurement ISQ (Osstell) value:<br>74.1 (at 12 mo)                                | : ISQ (Osstell) value:<br>74.1 (at 12 mo)  | Periotest value (Slemens): ISQ values at 12 mo: -5 ± 1.2 (at 6 mo) IL: 75.36 ± 5.88, CL: 75.64 ± 4.84  | ISQ values at 12 mo:<br>IL: 75.36 ± 5.88,<br>CL: 75.64 ± 4.84  | Periotest value:<br>-4 (at 360 d)  | ISQ value: 75 (at 6 mo)   | ÌSQ value:<br>71.7 ± 6.2 (at 6 mo)  | Not measured  |
| Time to definitive restoration   | 12 wk  | 24 wk  | IL: 1 wk, CL: 12 wk  | 6 wk   | 26 wk (for 39 implants),<br>NS (for the remaining<br>11 implants)   | NS  | 12 wk   |
| Use of antibiotics   | No antibiotics were prescribed   | Oral antibiotics 1 d before surgery and for 8 d afterward  | 2 g amoxicillin and clavulanic acid 2 h before surgery and for 4 d afterward   | SN   | 1 g amoxicillin 1 h before surgery and for 3 d afterward  | 1 g amoxicillin twice daily for 5 d   | 2 g amoxicillin 2 h before<br>surgery   |
| Follow-up period   | 36 mo  | 24 mo  | 12 mo  | 12 mo  | 6-12 mo   | 12 mo   | 12 mo   |
| Implant survival rate (%)  | 100  | 100  | 7, CL: 100   | 6.09   | 100   | 96.7  | IL: 93.3, CL: 100   |
| Quality category   | Average  | Average  | Better   | Average  | Average   | Average   | Better  |
|  |  | The second secon |  |  |   |   |   |

CL = conventional loading; RCT = randomized controlled trial; IL = immediate loading; IR = immediate restoration; ISQ = implant stability quotient; NS = not specified.

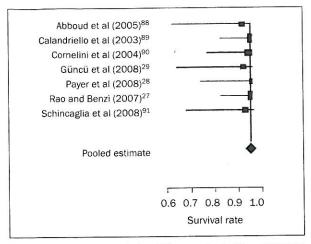


Fig 5 Forest plot analysis of the survival of immediately restored/loaded single implants in healed molar sites.

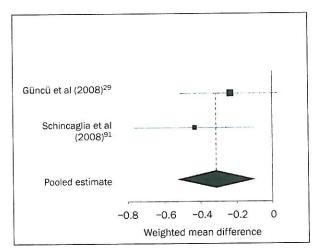


Fig 7 Forest plot analysis of immediate restoration/loading versus conventional loading in healed molar sites: Marginal bone changes after 12 months.

two groups at any point in time, whereas the other study reported significant differences in favor of the immediate loading group.<sup>91</sup>

In terms of implant failure, the meta-analysis found no differences in treatment effect between the immediate and conventional loading groups (relative risk of 3.0, 95% Cl: 0.33 to 27.16, P = .33, Fig 6). No statistically significant heterogeneity was detected between the studies (P = 1.0 with  $I^2 = 0\%$ ). With respect to marginal bone level changes, the meta-analysis revealed a significant difference between the two loading protocols, with a mean difference of -0.31 (95% Cl: -0.53 to -0.096, P = .005; Fig 7). Despite the inclusion of two trials comparing different loading protocols, no heterogeneity was found among the seven included studies (P = .37).

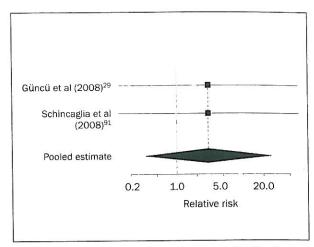


Fig 6 Forest plot analysis of immediate restoration/loading versus conventional loading in healed molar sites: Single implant survival rate.

### DISCUSSION

The guidelines of the AO SSID workshop<sup>45,46</sup> were followed in conducting this systematic review and meta-analysis. Two PICO questions were formulated. A total of 1,013 single implants placed in fresh molar extraction sites was included in the first analysis to investigate the overall estimate of the survival of immediately placed implants and the corresponding 95% CI. The second analysis included 188 immediately restored/loaded single implants placed into healed molar sites to establish an overall survival rate and its corresponding 95% CI. The pooling of the extracted data in both analyses showed high survival rates: 99.0% for immediate placement and 97.9% for immediate loading of single implants in healed molar sites.

In addition, the meta-analysis showed a statistically similar implant failure rate between immediate and delayed placement of single implants in molar extraction sockets. Likewise, no significant difference in implant failure was found between immediate and delayed implant loading with single crowns in healed molar sites. However, favorable marginal bone level changes were detected after immediate loading of single implants with crowns in healed molar ridges.

It is acknowledged that there are other extensive reviews in the literature evaluating immediate placement and loading.<sup>31–44</sup> Seven review articles<sup>31–37</sup> have specifically discussed the timing of implant placement. The earliest review<sup>31</sup> was a narrative literature review that included both experimental and human studies. It described the importance of placing implants beyond the apex of the tooth socket to achieve primary implant stability and highlighted the need for more investigations to evaluate the use of bone grafting and/or membranes for gap filling. Chen

et al<sup>32</sup> conducted a systematic review to study the histologic basis and clinical outcomes of immediate and delayed implant placement. The search was limited to the MEDLINE electronic database and the bibliographies of relevant articles. All types of study designs, including case series, were identified and included. Meta-analysis was not performed because of the heterogeneity of the included studies. However, comparable short-term survival rates of both immediate and delayed placement were observed. The authors recommended a soft tissue healing period of 4 to 8 weeks before placing implants.

In a Cochrane review,33 a more comprehensive search strategy was adopted to select all RCTs that examined immediate, immediate-delayed (early), and delayed placement in both partially and completely edentulous situations. Using data from only two RCTs, the reviewers showed that immediate and immediatedelayed placement may offer more advantages in terms of esthetics, alveolar bone height preservation, and patient satisfaction, but the report emphasized the need for more well-designed RCTs to evaluate long-term outcomes. In addition to the broad inclusion criteria, which may have affected the homogeneity of the studies and thus the validity of the meta-analytic results, this review was published in 2006 and has not yet been updated, as is recommended by the Cochrane Collaboration.92

As part of the AO workshop in 2006, Quirynen et al<sup>34</sup> published a comprehensive systematic review to study the time of implant placement. The review included all study designs investigating single-tooth and partially and completely edentulous situations. A classification for bone-implant gaps was presented and the recommendation made that bone defects > 2 mm lateral to implants be treated using bone grafts. This review showed an overall failure rate of 4% to 5% when implants were placed in extraction sockets. A higher failure rate was demonstrated with immediate loading of immediately placed implants, particularly for minimally rough-surfaced implants. In addition, the review called for strict inclusion criteria for immediate placement, including proper patient selection and treatment planning. Schropp and Isidor35 followed a search strategy that included PubMed and a manual search. They reported a summary of clinical guidelines for immediate or early implant placement and considered the combination of immediate implant placement and immediate restoration/loading a viable treatment option in the anterior mandible. However, these authors found limited literature to support these protocols in the maxilla and posterior mandible. Careful patient selection was recommended for replacement of molars, as this was considered a more challenging procedure. Further clinical

guidelines and a description of surgical technique for immediate implant placement were presented by Becker and Goldstein.<sup>36</sup> A more recent review<sup>37</sup> discussed in a narrative way the advantages and disadvantages of immediate placement and summarized the most important clinical criteria to achieve a successful outcome.

Specifically regarding loading protocols, seven review papers have also been published.38-44 Gapski et al<sup>38</sup> conducted a Medline search to critically review and analyze immediate restoration/loading protocols. Primary implant stability was considered the most important requirement for loading. This review discussed the influence of host-, implant-, and occlusion-related factors on the outcome of immediate implant loading. A need for-long term prospective studies to investigate the key factors in successful outcomes and the effect of systemic conditions on immediate loading protocols was emphasized. Attard and Zarb<sup>39</sup> performed an extensive literature review that included all types of studies evaluating immediate or early restoration/loading approaches in partially and completely edentulous patients. Based on 90 studies published between 1975 and 2004, the authors concluded that predictable outcomes can only be achieved in the anterior mandible because of a lack of evidence in the maxilla and posterior mandible. Further long-term studies for separate clinical situations were recommended to allow more meaningful comparisons. Nkenke and Fenner<sup>40</sup> analyzed 38 studies published between 1994 and 2005 and summed up the available literature for immediate restoration/loading in all clinical applications. The authors called for more long-term studies and concluded that there was still no evidence to recommend specific inclusion criteria, implant stability values, and bone quality for successful immediate restoration/loading. Jokstad and Carr<sup>41</sup> conducted a comprehensive systematic review as part of the AO workshop. The review included 22 studies published between 1990 and 2005. The authors showed that the literature lacked a proper methodology and mostly presented short-term results but concluded that immediate and early restoration/loading protocols can result in comparable outcomes to conventional protocols in many clinical situations. Moreover, an updated Cochrane review<sup>42</sup> suggested that immediate and early loading techniques can be realistic alternatives to conventional loading in selected situations. The meta-analysis included 11 RCTs that compared different loading protocols in all clinical situations. No statistically significant differences were found. Although the Cochrane Collaboration's guidelines follow a strict strategy in terms of quality assessment and data analysis, the differences between

partial and complete edentulism in terms of ridge dimensions, occlusal forces, and the presence of teeth may add to the heterogeneity of the included studies and thus compromise the validity of the meta-analytic results. In a more recent review, Henry and Liddelow<sup>43</sup> reviewed the biologic basis and guidelines for immediate loading protocols and presented case reports that included both partially and fully edentulous situations. Sennerby and Gottlow<sup>44</sup> reviewed six controlled studies published after 2005. The authors concluded that immediate and early restoration/ loading is a safe and viable alternative in selected cases. It is worth noting that this review revealed no significant differences between machined or moderately rough-surfaced implants following immediate loading. Again, more long-term RCTs were urged.

All the aforementioned review studies have assessed immediate placement and/or immediate restoration/loading of both partial and completely edentulous situations and single-tooth restorations. None of the reviews were limited to single implants or molar sites. The inclusion of such broad criteria made meta-analysis difficult because of the heterogeneity of the studies. In contrast, the present study is different from other previous reviews in many ways. First, the search strategy adopted an extensive approach that included several electronic databases; the Cochrane, United Kingdom, Australian, and New Zealand trials registries; conference proceedings; and abstracts. In addition, the search included a hand search of several journals as well as unpublished and non-English language publications. Second, the authors attempted to answer two focused clinical questions related to placement and loading of single implants in molar areas. Third, the inclusion criteria accepted studies of similar design and quality, permitting the use of a meta-analytic approach and providing answers with a high level of evidence.

However, the authors acknowledge that the present meta-analysis has several limitations. First, the quality of the existing literature was questionable, as most of the included studies were classified as fair or average. Second, the variability in study designs and follow-up periods may have contributed to the heterogeneity among studies. Nevertheless, adequate homogeneity was demonstrated by the close proximity of the boxes to the lines seen on the forest plots, in addition to the test of heterogeneity. Third, of the controlled studies, only three compared immediate placement with other placement protocols, and only two included both immediate and delayed loading groups. Therefore, the inclusion of such a small number of studies might be insufficient to draw solid conclusions. Fourth, few data were available regarding implants in molar sites with respect to other important peri-implant parameters, such as probing pocket depths, probing attachment level, and width of keratinized tissue in molar sites, as a result of limited available data. Finally, the authors' search strategy was not explicitly focused on studies that combined immediate placement of single implants in molar extraction sockets (both maxillary and mandibular) with immediate restoration/loading using a crown. The authors were unable to identify any published controlled clinical trials that compared this treatment option to conventional approaches. Thus, the authors chose to address the question indirectly by conducting simultaneous parallel searches for studies of immediate placement or immediate restoration of implants in molar extraction sockets.

The pooled estimate of the implant survival rate and hence the main findings of the current review showed that immediate placement of single implants in molar extraction sockets and immediate restoration/loading of single implants in healed molar sites are both predictable procedures, provided that strict selection criteria are followed. This review highlights the need for more randomized controlled studies, preferably with long-term outcomes, that include larger numbers of participants to provide further evidence for immediate placement of implants in molar extraction sockets or immediate restoration/loading of single implants in molar sites in healed sites.

### CONCLUSIONS

This systematic review and meta-analysis identified nine articles that met inclusion criteria; these demonstrated a high survival rate for single implants immediately placed in molar extraction sockets that was comparable to implant placement in healed sites. Survival rates were also high for immediate restoration/loading of single implants in healed molar sites. The literature currently lacks any controlled studies comparing the combined approach of immediate restoration/loading of immediately placed implants to more traditional implant protocols in molar regions.

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