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Efficiency, safety and complications of autologous fat grafting to the eyelids and periorbital area: A systematic review and meta-analysis

Fan Yang¹¹Department of Plastic Surgery and Burns, Tangdu Hospital, The Fourth Military Medical University, Shaanxi, China

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crystal yangfan

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ABSTRACT

Background: In recent years, autologous fat grafting (AFG), also known as fat transfer or lipofilling, has been widely performed for periorbital rejuvenation and defect correction, although the evidence regarding its efficacy and safety is still lacking. Besides, with respect to the periorbital region, it is invariably the earliest appearance area of the facial aging phenomenon. Therefore, a systematic review and meta-analysis is needed to evaluate the efficacy and safety of this technique.

Methods: A literature search was performed in PubMed, Embase, and the Cochrane library databases on November 11, 2020, adhering to the PRISMA guidelines, to identify all relevant articles. Then, a data extraction and standardization process was performed to assess all outcome data. Ultimately, the data were assessed using a random effects regression model with comprehensive meta-analysis software.

Results: Thirty-six studies consisting of four cohorts and 32 case series with a total of 3997 cases were included. The mean follow-up was 1.5 years. Meta-analysis revealed a relatively high satisfaction rate of 93.7% (95% CI, 89.6%–96.2%). Frequent complications in 3997 patients receiving AFG were edema, chemosis, and contour irregularity, with an overall complication rate of 7.7% (95% CI, 4.3%–13.3%).

Conclusion: This systematic review and meta-analysis showed that AFG for rejuvenation of eyelids and periorbital area provided a high satisfaction rate and did not result in severe complications. Therefore, AFG might be performed safely for periorbital rejuvenation and reconstruction.

THIS PROTOCOL ACCOMPANIES THE FOLLOWING PUBLICATION

References 1. Coleman SR. Longterm survival of fat transplants controlled demonstrations. *Aesthetic Plastic Surgery*. 1995;19:421-5. 2. Konn M. Structural fat grafting. *British Journal of Surgery*. 2005;92(5):657-. doi: 10.1002/bjs.4962. 3. Ramil ME. Fat Grafting in the Hollow Upper Eyelids and Volumetric Upper Blepharoplasty. *Plast Reconstr Surg*. 2017;140(5):889-97. Epub 2017/07/29. doi: 10.1097/PRS.0000000000003758. PubMed PMID: 28753160. 4. Lee W, Kwon SB, Oh SK, Yang EJ. Correction of sunken upper eyelid with orbital fat transposition flap and dermofat graft. *J Plast Reconstr Aesthet Surg*. 2017;70(12):1768-75. Epub 2017/06/18. doi: 10.1016/j.bjps.2017.05.003. PubMed PMID: 28619482. 5. Lin TM, Lin TY, Huang YH, Hsieh TY, Chou CK, Takahashi H, et al. Fat Grafting for Recontouring Sunken Upper Eyelids With Multiple Folds in Asians-Novel Mechanism for Neoformation of Double Eyelid Crease. *Annals of plastic surgery*. 2016;76(4):371-5. Epub 2015/12/19. doi: 10.1097/sap.0000000000000668. PubMed PMID: 26678103. 6. Lin TM, Lin TY, Chou CK, Lai CS, Lin SD. Application of microautologous fat transplantation in the correction of sunken upper eyelid. *Plast Reconstr Surg Glob Open*. 2014;2(11):e259. Epub 2014/12/17. doi: 10.1097/GOX.0000000000000141. PubMed PMID: 25506542; PubMed Central PMCID: PMC4255902. 7. Yoonho Lee MD, Ph.D., Sungtack Kwon, M.D., Ph.D., and Kun Hwang, M.D., Ph.D. Correction of Sunken and/or Multiply Folded Upper Eyelid by Fascia-Fat Graft. *Plast Reconstr Surg*. 2001;107(1):15-9. 8. Tonnard PL, Verpaele AM, Zeltzer AA. Augmentation blepharoplasty: a review of 500 consecutive patients. *Aesthetic surgery journal*. 2013;33(3):341-52. Epub 2013/03/22. doi: 10.1177/1090820x13478966. PubMed PMID: 23515379. 9. Rehman J, Traktuev D, Li J, Merfeld-Clauss S, Temm-Grove CJ, Bovenkerk JE, et al. Secretion of angiogenic and antiapoptotic factors by human adipose stromal cells. *Circulation*. 2004;109(10):1292-8. Epub 2004/03/03. doi: 10.1161/01.CIR.0000121425.42966.F1. PubMed PMID: 14993122. 10. Varghese J, Griffin M, Mosahebi A, Butler P. Systematic review of patient factors affecting adipose stem cell viability and function: implications for regenerative therapy. *Stem Cell Res Ther*. 2017;8(1):45. Epub 2017/03/01. doi:

10.1186/s13287-017-0483-8. PubMed PMID: 28241882; PubMed Central PMCID: PMC5329955. 11. Trepsat F. Periorbital rejuvenation combining fat grafting and blepharoplasties. *Aesthetic Plast Surg.* 2003;27(4):243-53. Epub 2004/04/03. doi: 10.1007/s00266-003-2126-y. PubMed PMID: 15058544. 12. Boureaux E, Chaput B, Bannani S, Herlin C, De Runz A, Carloni R, et al. Eyelid fat grafting: Indications, operative technique and complications; a systematic review. *Journal of cranio-maxillo-facial surgery : official publication of the European Association for Cranio-Maxillo-Facial Surgery.* 2016;44(4):374-80. Epub 2016/02/18. doi: 10.1016/j.jcms.2015.12.013. PubMed PMID: 26880013. 13. Maamari RN, Massry GG, Holds JB. Complications Associated with Fat Grafting to the Lower Eyelid. *Facial plastic surgery clinics of North America.* 2019;27(4):435-41. Epub 2019/10/08. doi: 10.1016/j.fsc.2019.07.001. PubMed PMID: 31587763. 14. Shue S, Kurlander DE, Guyuron B. Fat Injection: A Systematic Review of Injection Volumes by Facial Subunit. *Aesthetic plastic surgery.* 2018;42(5):1261-70. Epub 2017/08/10. doi: 10.1007/s00266-017-0936-6. PubMed PMID: 28791455. 15. Cetinkaya A, Devoto MH. Periocular fat grafting: indications and techniques. *Curr Opin Ophthalmol.* 2013;24(5):494-9. Epub 2013/08/09. doi: 10.1097/ICU.0b013e3283634841. PubMed PMID: 23925063. 16. Knobloch K, Yoon U, Vogt PM. Preferred reporting items for systematic reviews and meta-analyses (PRISMA) statement and publication bias. *J Craniomaxillofac Surg.* 2011;39(2):91-2. Epub 2010/12/15. doi: 10.1016/j.jcms.2010.11.001. PubMed PMID: 21145753. 17. Pelle-Ceravolo M, Angelini M. Properly Diluted Fat (PDF): An Easy and Safe Approach to Periocular Fat Grafting. *Aesthetic surgery journal.* 2019. Epub 2019/02/13. doi: 10.1093/asj/sjz039. PubMed PMID: 30753276. 18. Essuman VA, Tagoe NN, Ndanu TA, Ntim-Amponsah CT. Dermis-fat grafts and enucleation in Ghanaian children: 5 years' experience. *Ghana Med J.* 2014;48(4):204-7. Epub 2015/02/25. doi: 10.4314/gmj.v48i4.6. PubMed PMID: 25709135; PubMed Central PMCID: PMC4335434. 19. Jason D. Meier MRAG, MD; Mark J. Glasgold, MD. Autologous Fat Grafting Long-term Evidence of Its Efficacy in Midfacial Rejuvenation. 20. Bernardini FP, Gennai A, Izzo L, Zambelli A, Repaci E, Baldelli I, et al. Superficial Enhanced Fluid Fat Injection (SEFFI) to Correct Volume Defects and Skin Aging of the Face and Periocular Region. *Aesthetic surgery journal.* 2015;35(5):504-15. Epub 2015/04/26. doi: 10.1093/asj/sjv001. PubMed PMID: 25911629. 21. Kim HS, Choi CW, Kim BR, Youn SW. Effectiveness of Transconjunctival Fat Removal and Resected Fat Grafting for Lower Eye Bag and Tear Trough Deformity. *JAMA Facial Plast Surg.* 2019;21(2):118-24. Epub 2018/11/13. doi: 10.1001/jamafacial.2018.1307. PubMed PMID: 30418468; PubMed Central PMCID: PMC6439811. 22. Huang SH, Lin YN, Lee SS, Huang YH, Takahashi H, Chou CK, et al. Three Simple Steps for Refining Transcutaneous Lower Blepharoplasty for Aging Eyelids: The Indispensability of Micro-Autologous Fat Transplantation. *Aesthet Surg J.* 2019;39(11):1163-77. Epub 2019/01/23. doi: 10.1093/asj/sjz005. PubMed PMID: 30668643. 23. Kim J, Shin H, Lee M, Shin D, Kim S, Jo D, et al. Percutaneous Autologous Fat Injection Following 2-Layer Flap Lower Blepharoplasty for the Correction of Tear Trough Deformity. *The Journal of craniofacial surgery.* 2018;29(5):1241-4. Epub 2018/04/03. doi: 10.1097/scs.0000000000004552. PubMed PMID: 29608475. 24. Chen H, Zhang Q, Qiu Q, Yang Z. Autologous Fat Graft for the Treatment of Sighted Posttraumatic Enophthalmos and Sunken Upper Eyelid. *Ophthalmic Plast Reconstr Surg.* 2018;34(4):381-6. Epub 2018/01/26. doi: 10.1097/IOP.0000000000001028. PubMed PMID: 29369151. 25. Miranda SG, Codner MA. Micro Free Orbital Fat Grafts to the Tear Trough Deformity during Lower Blepharoplasty. *Plast Reconstr Surg.* 2017;139(6):1335-43. Epub 2017/02/16. doi: 10.1097/PRS.0000000000003356. PubMed PMID: 28198772. 26. Ma Z, Chen G, Huang J, Wang J, Huang W. Clinical application of orbital fat autograft for tear trough deformity through tranconjunctival approach. *Chin J Plast Surg.* 2017;33(2):116-9. 27. Gennai A, Zambelli A, Repaci E, Quarto R, Baldelli I, Fraternali G, et al. Skin Rejuvenation and Volume Enhancement with the Micro Superficial Enhanced Fluid Fat Injection (M-SEFFI) for Skin Aging of the Periocular and Perioral Regions. *Aesthetic surgery journal.* 2017;37(1):14-23. Epub 2016/06/01. doi: 10.1093/asj/sjw084. PubMed PMID: 27241362. 28. Chiu CY, Shen YC, Zhao QF, Hong FL, Xu JH. Treatment of Tear Trough Deformity: Fat Repositioning versus Autologous Fat Grafting. *Aesthetic plastic surgery.* 2017;41(1):73-80. Epub 2016/12/23. doi: 10.1007/s00266-016-0692-z. PubMed PMID: 28008460. 29. Jiang J, Wang X, Chen R, Xia X, Sun S, Hu K. Tear trough deformity: different types of anatomy and treatment options. *Postepy dermatologii i alergologii.* 2016;33(4):303-8. Epub 2016/09/09. doi: 10.5114/ada.2016.61607. PubMed PMID: 27605904; PubMed Central PMCID: PMC5004220. 30. Park S, Kim B, Shin Y. Correction of superior sulcus deformity with orbital fat anatomic repositioning and fat graft applied to retro-orbicularis oculi fat for Asian eyelids. *Aesthetic plastic surgery.* 2011;35(2):162-70. Epub 2010/09/14. doi: 10.1007/s00266-010-9574-y. PubMed PMID: 20835821. 31. Litwin AS, Poitelea C, Tan P, Ziahosseini K, Malhotra R. Complications and outcomes of grafting of posterior orbital fat into the lower lid-cheek junction during orbital decompression. *Orbit (Amsterdam, Netherlands).* 2018;37(2):128-34. Epub 2017/10/13. doi: 10.1080/01676830.2017.1383452. PubMed PMID: 29023175. 32. Chang HS, Lee D, Taban M, Douglas RS, Goldberg RA. "En-glove" lysis of lower eyelid retractors with AlloDerm and dermis-fat grafts in lower eyelid

retraction surgery. *Ophthalmic Plast Reconstr Surg*. 2011;27(2):137-41. Epub 2010/06/22. doi: 10.1097/IOP.0b013e3181c53d38. PubMed PMID: 20562664. 33. Çoban Karataş M, Yaycıoğlu RA, Canan H. Orbital Dermis-Fat Graft Transplantation: Results in Primary and Secondary Implantation. *Türk Oftalmoloji Dergisi*. 2015;45(2):65-70. doi: 10.4274/tjo.55823. 34. Roh MR, Kim TK, Chung KY. Treatment of infraorbital dark circles by autologous fat transplantation: a pilot study. *The British journal of dermatology*. 2009;160(5):1022-5. Epub 2009/05/13. doi: 10.1111/j.1365-2133.2009.09066.x. PubMed PMID: 19434788. 35. Youn S, Shin JI, Kim JD, Kim JT, Kim YH. Correction of infraorbital dark circles using collagenase-digested fat cell grafts. *Dermatologic surgery : official publication for American Society for Dermatologic Surgery [et al]*. 2013;39(5):766-72. Epub 2013/02/27. doi: 10.1111/dsu.12140. PubMed PMID: 23437990. 36. Pelle-Ceravolo M, Angelini M. Properly Diluted Fat (PDF): An Easy and Safe Approach to Periocular Fat Grafting. *Aesthetic surgery journal*. 2020;40(1):19-33. Epub 2019/02/13. doi: 10.1093/asj/sjz039. PubMed PMID: 30753276. 37. Park JY, Kim N. Periorbital Lipogranuloma after Facial Autologous Fat Injection and Its Treatment Outcomes. *Korean journal of ophthalmology : KJO*. 2016;30(1):10-6. Epub 2016/02/13. doi: 10.3341/kjo.2016.30.1.10. PubMed PMID: 26865798; PubMed Central PMCID: PMC4742640. 38. Li XQ, Wang TL, Wang JQ. Ptosis: An Underestimated Complication after Autologous Fat Injection into the Upper Eyelid. *Aesthetic surgery journal*. 2015;35(6):NP147-53. Epub 2015/08/01. doi: 10.1093/asj/sjv015. PubMed PMID: 26229134. 39. Rohrich RJ, Mahedia M, Shah N, Afroz P, Vishvanath L, Gupta RK. Role of Fractionated Fat in Blending the Lid-Cheek Junction. *Plast Reconstr Surg*. 2018;142(1):56-65. Epub 2018/06/08. doi: 10.1097/PRS.0000000000004526. PubMed PMID: 29878987. 40. Hsu VM, Stransky CA, Bucky LP, Percec I. Fat grafting's past, present, and future: why adipose tissue is emerging as a critical link to the advancement of regenerative medicine. *Aesthet Surg J*. 2012;32(7):892-9. Epub 2012/09/04. doi: 10.1177/1090820X12455658. PubMed PMID: 22942117. 41. Behr B, Tang C, Germann G, Longaker MT, Quarto N. Locally applied vascular endothelial growth factor A increases the osteogenic healing capacity of human adipose-derived stem cells by promoting osteogenic and endothelial differentiation. *Stem Cells*. 2011;29(2):286-96. Epub 2011/07/07. doi: 10.1002/stem.581. PubMed PMID: 21732486; PubMed Central PMCID: PMC3400547. 42. Tonnard P, Verpaale A, Peeters G, Hamdi M, Cornelissen M, Declercq H. Nanofat grafting: basic research and clinical applications. *Plastic and reconstructive surgery*. 2013;132(4):1017-26. Epub 2013/06/21. doi: 10.1097/PRS.0b013e31829fe1b0. PubMed PMID: 23783059. 43. Jiang S, Quan Y, Wang J, Cai J, Lu F. Fat Grafting for Facial Rejuvenation Using Stromal Vascular Fraction Gel Injection. *Clin Plast Surg*. 2020;47(1):73-9. Epub 2019/11/20. doi: 10.1016/j.cps.2019.09.001. PubMed PMID: 31739900. 44. Ho Quoc C, Taupin T, Guerin N, Delay E. Volumetric evaluation of fat resorption after breast lipofilling. *Ann Chir Plast Esthet*. 2015;60(6):495-9. Epub 2015/08/01. doi: 10.1016/j.anplas.2015.06.011. PubMed PMID: 26229038. 45. Stig-Frederik Trojahn Kølbe AF-N, Anders Bruun Mathiasen, Jens Jørgen Elberg, Roberto S Oliveri, Peter V Glovinski, Jens Kastrup, Maria Kirchhoff, Bo Sonnich Rasmussen, Maj-Lis Møller Talman, Carsten Thomsen, Ebbe Dickmeiss, Krzysztof Tadeusz Drzewiecki. Enrichment of autologous fat grafts with ex-vivo expanded adipose tissue-derived stem cells for graft survival: a randomised placebo-controlled trial. *Lancet*. 2013;382:1113-20. 46. Gir P, Brown SA, Oni G, Kashefi N, Mojallal A, Rohrich RJ. Fat grafting: evidence-based review on autologous fat harvesting, processing, reinjection, and storage. *Plast Reconstr Surg*. 2012;130(1):249-58. Epub 2012/06/30. doi: 10.1097/PRS.0b013e318254b4d3. PubMed PMID: 22743888. 47. Pusic AL, Klassen AF, Scott AM, Klok JA, Cordeiro PG, Cano SJ. Development of a new patient-reported outcome measure for breast surgery: the BREAST-Q. *Plast Reconstr Surg*. 2009;124(2):345-53. Epub 2009/08/01. doi: 10.1097/PRS.0b013e3181aee807. PubMed PMID: 19644246.

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References 1. Coleman SR. Longterm survival of fat transplants controlled demonstrations. *Aesthetic Plastic Surgery*. 1995;19:421-5. 2. Konn M. Structural fat grafting. *British Journal of Surgery*. 2005;92(5):657-. doi: 10.1002/bjs.4962. 3. Ramil ME. Fat Grafting in the Hollow Upper Eyelids and

Volumetric Upper Blepharoplasty. *Plast Reconstr Surg.* 2017;140(5):889-97. Epub 2017/07/29. doi: 10.1097/PRS.0000000000003758. PubMed PMID: 28753160. 4. Lee W, Kwon SB, Oh SK, Yang EJ. Correction of sunken upper eyelid with orbital fat transposition flap and dermofat graft. *J Plast Reconstr Aesthet Surg.* 2017;70(12):1768-75. Epub 2017/06/18. doi: 10.1016/j.bjps.2017.05.003. PubMed PMID: 28619482. 5. Lin TM, Lin TY, Huang YH, Hsieh TY, Chou CK, Takahashi H, et al. Fat Grafting for Recontouring Sunken Upper Eyelids With Multiple Folds in Asians-Novel Mechanism for Neoformation of Double Eyelid Crease. *Annals of plastic surgery.* 2016;76(4):371-5. Epub 2015/12/19. doi: 10.1097/sap.0000000000000668. PubMed PMID: 26678103. 6. Lin TM, Lin TY, Chou CK, Lai CS, Lin SD. Application of microautologous fat transplantation in the correction of sunken upper eyelid. *Plast Reconstr Surg Glob Open.* 2014;2(11):e259. Epub 2014/12/17. doi: 10.1097/GOX.0000000000000141. PubMed PMID: 25506542; PubMed Central PMCID: PMC4255902. 7. Yoonho Lee MD, Ph.D., Sungtaek Kwon, M.D., Ph.D., and Kun Hwang, M.D., Ph.D. Correction of Sunken and/or Multiply Folded Upper Eyelid by Fascia-Fat Graft. *Plast Reconstr Surg.* 2001;107(1):15-9. 8. Tonnard PL, Verpaele AM, Zeltzer AA. Augmentation blepharoplasty: a review of 500 consecutive patients. *Aesthetic surgery journal.* 2013;33(3):341-52. Epub 2013/03/22. doi: 10.1177/1090820x13478966. PubMed PMID: 23515379. 9. Rehman J, Traktuev D, Li J, Merfeld-Clauss S, Temm-Grove CJ, Bovenkerk JE, et al. Secretion of angiogenic and antiapoptotic factors by human adipose stromal cells. *Circulation.* 2004;109(10):1292-8. Epub 2004/03/03. doi: 10.1161/01.CIR.0000121425.42966.F1. PubMed PMID: 14993122. 10. Varghese J, Griffin M, Mosahebi A, Butler P. Systematic review of patient factors affecting adipose stem cell viability and function: implications for regenerative therapy. *Stem Cell Res Ther.* 2017;8(1):45. Epub 2017/03/01. doi: 10.1186/s13287-017-0483-8. PubMed PMID: 28241882; PubMed Central PMCID: PMC5329955. 11. Trepsat F. Periorbital rejuvenation combining fat grafting and blepharoplasties. *Aesthetic Plast Surg.* 2003;27(4):243-53. Epub 2004/04/03. doi: 10.1007/s00266-003-2126-y. PubMed PMID: 15058544. 12. Boureaux E, Chaput B, Bannani S, Herlin C, De Runz A, Carloni R, et al. Eyelid fat grafting: Indications, operative technique and complications; a systematic review. *Journal of cranio-maxillo-facial surgery : official publication of the European Association for Cranio-Maxillo-Facial Surgery.* 2016;44(4):374-80. Epub 2016/02/18. doi: 10.1016/j.jcms.2015.12.013. PubMed PMID: 26880013. 13. Maamari RN, Massry GG, Holds JB. Complications Associated with Fat Grafting to the Lower Eyelid. *Facial plastic surgery clinics of North America.* 2019;27(4):435-41. Epub 2019/10/08. doi: 10.1016/j.fsc.2019.07.001. PubMed PMID: 31587763. 14. Shue S, Kurlander DE, Guyuron B. Fat Injection: A Systematic Review of Injection Volumes by Facial Subunit. *Aesthetic plastic surgery.* 2018;42(5):1261-70. Epub 2017/08/10. doi: 10.1007/s00266-017-0936-6. PubMed PMID: 28791455. 15. Cetinkaya A, Devoto MH. Periocular fat grafting: indications and techniques. *Curr Opin Ophthalmol.* 2013;24(5):494-9. Epub 2013/08/09. doi: 10.1097/ICU.0b013e3283634841. PubMed PMID: 23925063. 16. Knobloch K, Yoon U, Vogt PM. Preferred reporting items for systematic reviews and meta-analyses (PRISMA) statement and publication bias. *J Craniomaxillofac Surg.* 2011;39(2):91-2. Epub 2010/12/15. doi: 10.1016/j.jcms.2010.11.001. PubMed PMID: 21145753. 17. Pelle-Ceravolo M, Angelini M. Properly Diluted Fat (PDF): An Easy and Safe Approach to Periocular Fat Grafting. *Aesthetic surgery journal.* 2019. Epub 2019/02/13. doi: 10.1093/asj/sjz039. PubMed PMID: 30753276. 18. Essuman VA, Tagoe NN, Ndanu TA, Ntim-Amponsah CT. Dermis-fat grafts and enucleation in Ghanaian children: 5 years' experience. *Ghana Med J.* 2014;48(4):204-7. Epub 2015/02/25. doi: 10.4314/gmj.v48i4.6. PubMed PMID: 25709135; PubMed Central PMCID: PMC4335434. 19. Jason D. Meier MRAG, MD; Mark J. Glasgold, MD. Autologous Fat Grafting Long-term Evidence of Its Efficacy in Midfacial Rejuvenation. 20. Bernardini FP, Gennai A, Izzo L, Zambelli A, Repaci E, Baldelli I, et al. Superficial Enhanced Fluid Fat Injection (SEFFI) to Correct Volume Defects and Skin Aging of the Face and Periocular Region. *Aesthetic surgery journal.* 2015;35(5):504-15. Epub 2015/04/26. doi: 10.1093/asj/sjv001. PubMed PMID: 25911629. 21. Kim HS, Choi CW, Kim BR, Youn SW. Effectiveness of Transconjunctival Fat Removal and Resected Fat Grafting for Lower Eye Bag and Tear Trough Deformity. *JAMA Facial Plast Surg.* 2019;21(2):118-24. Epub 2018/11/13. doi: 10.1001/jamafacial.2018.1307. PubMed PMID: 30418468; PubMed Central PMCID: PMC6439811. 22. Huang SH, Lin YN, Lee SS, Huang YH, Takahashi H, Chou CK, et al. Three Simple Steps for Refining Transcutaneous Lower Blepharoplasty for Aging Eyelids: The Indispensability of Micro-Autologous Fat Transplantation. *Aesthet Surg J.* 2019;39(11):1163-77. Epub 2019/01/23. doi: 10.1093/asj/sjz005. PubMed PMID: 30668643. 23. Kim J, Shin H, Lee M, Shin D, Kim S, Jo D, et al. Percutaneous Autologous Fat Injection Following 2-Layer Flap Lower Blepharoplasty for the Correction of Tear Trough Deformity. *The Journal of craniofacial surgery.* 2018;29(5):1241-4. Epub 2018/04/03. doi: 10.1097/scs.0000000000004552. PubMed PMID: 29608475. 24. Chen H, Zhang Q, Qiu Q, Yang Z. Autologous Fat Graft for the Treatment of Sighted Posttraumatic Enophthalmos and Sunken Upper Eyelid. *Ophthalmic Plast Reconstr Surg.* 2018;34(4):381-6. Epub 2018/01/26. doi: 10.1097/IOP.0000000000001028. PubMed PMID: 29369151. 25. Miranda SG, Codner

MA. Micro Free Orbital Fat Grafts to the Tear Trough Deformity during Lower Blepharoplasty. *Plast Reconstr Surg*. 2017;139(6):1335-43. Epub 2017/02/16. doi: 10.1097/PRS.0000000000003356. PubMed PMID: 28198772. 26. Ma Z, Chen G, Huang J, Wang J, Huang W. Clinical application of orbital fat autograft for tear trough deformity through tranconjunctival approach. *Chin J Plast Surg*. 2017;33(2):116-9. 27. Gennai A, Zambelli A, Repaci E, Quarto R, Baldelli I, Fraternali G, et al. Skin Rejuvenation and Volume Enhancement with the Micro Superficial Enhanced Fluid Fat Injection (M-SEFFI) for Skin Aging of the Periocular and Perioral Regions. *Aesthetic surgery journal*. 2017;37(1):14-23. Epub 2016/06/01. doi: 10.1093/asj/sjw084. PubMed PMID: 27241362. 28. Chiu CY, Shen YC, Zhao QF, Hong FL, Xu JH. Treatment of Tear Trough Deformity: Fat Repositioning versus Autologous Fat Grafting. *Aesthetic plastic surgery*. 2017;41(1):73-80. Epub 2016/12/23. doi: 10.1007/s00266-016-0692-z. PubMed PMID: 28008460. 29. Jiang J, Wang X, Chen R, Xia X, Sun S, Hu K. Tear trough deformity: different types of anatomy and treatment options. *Postepy dermatologii i alergologii*. 2016;33(4):303-8. Epub 2016/09/09. doi: 10.5114/ada.2016.61607. PubMed PMID: 27605904; PubMed Central PMCID: PMC45004220. 30. Park S, Kim B, Shin Y. Correction of superior sulcus deformity with orbital fat anatomic repositioning and fat graft applied to retro-orbicularis oculi fat for Asian eyelids. *Aesthetic plastic surgery*. 2011;35(2):162-70. Epub 2010/09/14. doi: 10.1007/s00266-010-9574-y. PubMed PMID: 20835821. 31. Litwin AS, Poitelea C, Tan P, Ziahosseini K, Malhotra R. Complications and outcomes of grafting of posterior orbital fat into the lower lid-cheek junction during orbital decompression. *Orbit (Amsterdam, Netherlands)*. 2018;37(2):128-34. Epub 2017/10/13. doi: 10.1080/01676830.2017.1383452. PubMed PMID: 29023175. 32. Chang HS, Lee D, Taban M, Douglas RS, Goldberg RA. "En-glove" lysis of lower eyelid retractors with AlloDerm and dermis-fat grafts in lower eyelid retraction surgery. *Ophthalmic Plast Reconstr Surg*. 2011;27(2):137-41. Epub 2010/06/22. doi: 10.1097/IOP.0b013e3181c53d38. PubMed PMID: 20562664. 33. Çoban Karataş M, Yaycıoğlu RA, Canan H. Orbital Dermis-Fat Graft Transplantation: Results in Primary and Secondary Implantation. *Türk Oftalmoloji Dergisi*. 2015;45(2):65-70. doi: 10.4274/tjo.55823. 34. Roh MR, Kim TK, Chung KY. Treatment of infraorbital dark circles by autologous fat transplantation: a pilot study. *The British journal of dermatology*. 2009;160(5):1022-5. Epub 2009/05/13. doi: 10.1111/j.1365-2133.2009.09066.x. PubMed PMID: 19434788. 35. Youn S, Shin JI, Kim JD, Kim JT, Kim YH. Correction of infraorbital dark circles using collagenase-digested fat cell grafts. *Dermatologic surgery : official publication for American Society for Dermatologic Surgery [et al]*. 2013;39(5):766-72. Epub 2013/02/27. doi: 10.1111/dsu.12140. PubMed PMID: 23437990. 36. Pelle-Ceravolo M, Angelini M. Properly Diluted Fat (PDF): An Easy and Safe Approach to Periocular Fat Grafting. *Aesthetic surgery journal*. 2020;40(1):19-33. Epub 2019/02/13. doi: 10.1093/asj/sjz039. PubMed PMID: 30753276. 37. Park JY, Kim N. Periorbital Lipogranuloma after Facial Autologous Fat Injection and Its Treatment Outcomes. *Korean journal of ophthalmology : KJO*. 2016;30(1):10-6. Epub 2016/02/13. doi: 10.3341/kjo.2016.30.1.10. PubMed PMID: 26865798; PubMed Central PMCID: PMC4742640. 38. Li XQ, Wang TL, Wang JQ. Ptosis: An Underestimated Complication after Autologous Fat Injection into the Upper Eyelid. *Aesthetic surgery journal*. 2015;35(6):Np147-53. Epub 2015/08/01. doi: 10.1093/asj/sjv015. PubMed PMID: 26229134. 39. Rohrich RJ, Mahedia M, Shah N, Afroz P, Vishvanath L, Gupta RK. Role of Fractionated Fat in Blending the Lid-Cheek Junction. *Plast Reconstr Surg*. 2018;142(1):56-65. Epub 2018/06/08. doi: 10.1097/PRS.0000000000004526. PubMed PMID: 29878987. 40. Hsu VM, Stransky CA, Bucky LP, Percec I. Fat grafting's past, present, and future: why adipose tissue is emerging as a critical link to the advancement of regenerative medicine. *Aesthet Surg J*. 2012;32(7):892-9. Epub 2012/09/04. doi: 10.1177/1090820X12455658. PubMed PMID: 22942117. 41. Behr B, Tang C, Germann G, Longaker MT, Quarto N. Locally applied vascular endothelial growth factor A increases the osteogenic healing capacity of human adipose-derived stem cells by promoting osteogenic and endothelial differentiation. *Stem Cells*. 2011;29(2):286-96. Epub 2011/07/07. doi: 10.1002/stem.581. PubMed PMID: 21732486; PubMed Central PMCID: PMC3400547. 42. Tonnard P, Verpaele A, Peeters G, Hamdi M, Cornelissen M, Declercq H. Nanofat grafting: basic research and clinical applications. *Plastic and reconstructive surgery*. 2013;132(4):1017-26. Epub 2013/06/21. doi: 10.1097/PRS.0b013e31829fe1b0. PubMed PMID: 23783059. 43. Jiang S, Quan Y, Wang J, Cai J, Lu F. Fat Grafting for Facial Rejuvenation Using Stromal Vascular Fraction Gel Injection. *Clin Plast Surg*. 2020;47(1):73-9. Epub 2019/11/20. doi: 10.1016/j.cps.2019.09.001. PubMed PMID: 31739900. 44. Ho Quoc C, Taupin T, Guerin N, Delay E. Volumetric evaluation of fat resorption after breast lipofilling. *Ann Chir Plast Esthet*. 2015;60(6):495-9. Epub 2015/08/01. doi: 10.1016/j.anplas.2015.06.011. PubMed PMID: 26229038. 45. Stig-Frederik Trojahn Kølbe AF-N, Anders Bruun Mathiasen, Jens Jørgen Elberg, Roberto S Oliveri, Peter V Glovinski, Jens Kastrup, Maria Kirchhoff, Bo Sonnich Rasmussen, Maj-Lis Møller Talman, Carsten Thomsen, Ebbe Dickmeiss, Krzysztof Tadeusz Drzewiecki. Enrichment of autologous fat grafts with ex-vivo expanded adipose tissue-derived stem cells for graft survival: a randomised placebo-controlled trial. *Lancet*. 2013;382:1113-20. 46. Gir P, Brown SA, Oni G, Kashefi N, Mojallal A, Rohrich RJ. Fat grafting:

evidence-based review on autologous fat harvesting, processing, reinjection, and storage. *Plast Reconstr Surg*. 2012;130(1):249-58. Epub 2012/06/30. doi: 10.1097/PRS.0b013e318254b4d3. PubMed PMID: 22743888. 47. Pusic AL, Klassen AF, Scott AM, Klok JA, Cordeiro PG, Cano SJ. Development of a new patient-reported outcome measure for breast surgery: the BREAST-Q. *Plast Reconstr Surg*. 2009;124(2):345-53. Epub 2009/08/01. doi: 10.1097/PRS.0b013e3181aee807. PubMed PMID: 19644246.

KEYWORDS

Eyelid, Periorbital area, Fat grafting, Adipose-derived stem cell, Facial rejuvenation

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MATERIALS TEXT

Search strategy

The research objectives were to identify, assess, and synthesize the evidence examining the efficacy and safety of AFG in the periorbital area. This review was performed in accordance with the PRISMA guidelines. This comprehensive, reproducible, and electronic search was performed via the combination of PubMed, Embase, and Cochrane Library databases. The following keywords were used: [("fat grafting" OR "lipograft" OR "lipoinjection" OR "lipotransfer" OR "fat transfer" OR "fat transplant" OR "lipostructure" OR "lipofilling" OR "fat injection" OR "lipomodeling" OR "fat transplantation") AND ("eyelid" OR "periorbital")]. A systematic database search was carried out before November 11, 2020. There were no restrictions with respect to language.

Inclusion and exclusion criteria

Selected studies met the following criteria: (1) clinical trials of all designs, from the highest level of evidence from randomized trials (if available) to prospective or retrospective observational studies (case series: at least five cases) involving patients receiving AFG for periorbital rejuvenation and reconstruction; (2) the treatment used for periorbital rejuvenation and reconstruction was stated clearly; (3) the study stated the concrete data of postoperative effects; (4) studies with complete follow-up (at least 3 months). Exclusion criteria were as follows: (1) patients with a history of other eyelid surgery or treatment; (2) reviews, letters, commentaries, reply, discussion, and so on; (3) studies with incomplete or ambiguous or overlapped data; (4) studies not related to the objective of this review.

Data collection

Two independent reviewers scrutinized the titles, abstracts, and full text of the retrieved articles. If there was any disagreement between the two reviewers, another independent investigator was consulted to reach a consensus. Moreover, a blinded method was used to ensure quality. Data extracted from the eligible articles included the following parts: authors, date of publication, place of study, number of patients, ages of patients, indications, AFG techniques, follow-up time, study design, evidence level, complications, anesthetic evaluation, and satisfaction rates. Then a data extraction sheet was set in Excel (Microsoft, Redmond, Washington, USA). Additionally, each article was assessed for the risk of bias in accordance with the methodological standards listed in the non-comparative case series checklist (for case series) and Ottawa-Newcastle Scale (for cohort studies), respectively (<https://www.ncbi.nlm.nih.gov/books/NBK35156/>) (Table S1, Table S2).

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- 1 We will search the PubMed, Cochrane library, and Embase databases for articles published prior to November 11, 2020.
- 2 We will scrutinize the titles, abstracts, and full text of the retrieved articles and screen out articles that do not meet the inclusion and exclusion criteria.
- 3 We will perform meta-analysis by a comprehensive meta-analysis software (CMA), version 2.2.050 (Biostat, Englewood, NJ, USA). A random-effects meta-analysis will be conducted to analyze all the evidence.