

MANAGEMENT OF VARICOCELE IN ADOLESCENCE PATIENTS

Ali ATAN¹

Alparslan KAPISIZ²

The aim of varicocele treatment in adolescents is to keep gonadal functions and semen parameters to the maximum. As a result, it is aimed not to impair fertility in adulthood (1). Management and treatment of varicocele in childhood and adolescence are poorly defined and still debated as adolescents with varicocele are highly heterogeneous, due to rapid changes in hormone levels and the stage of pubertal development (2). Which of the adolescents with varicocele should be treated, when to start the treatment and which treatment method to choose are still a controversial issues. Firstly, adolescent and his family should be informed in detail about varicocele and surgical treatment by the pediatric urologist.

TREATMENT INDICATIONS

As the indications for varicocelectomy in adolescence are not clear, this makes a standard approach more difficult to these patients. Difference in right and left testis sizes, scrotal pain, hormonal values and, if possible, semen analysis parameters are considered important factors that can be used in the decision of varicocelectomy (3). However, the data regarding these parameters are sometimes conflicting. Testicular hypotrophy or testicular growth arrest is mostly considered the first indication to correct varicocele in pediatric age (4-7). In last EAU

¹ Prof. MD, Gazi University Faculty of Medicine, Department of Urology, aliatanpitt@hotmail.com, ORCID iD: 0000-0002-7114-068X

² Asist. Prof.MD, Gazi University Faculty of Medicine, Department of Pediatric Surgery, alparslankapisiz@gazi.edu.tr, ORCID iD: 0000-0002-4803-8900

Table 1. Varicocele surgical techniques: pros and cons (30).

Tecnique	Pros	Cons
Open retroperitoneal high ligation (Palomo)	Complete ligation	General anesthesia, Higher hydrocele risk
Microsurgical lymphatic sparing Palomo	Complete ligation Lower hydrocele risk Less invasive (local anesthesia)	Access to operating microscope
Microsurgical subinguinal or inguinal surgery	Lower recurrence rate Lower hydrocele risk	Access to operating microscope
Laparoscopic surgery	Bilateral varicocele Higher magnification Lower recurrence rate	High cost More invasive (intraperitoneal) General anesthesia
Sclero-embolization	Minimally invasive Short time Outpatient	Limited applicability Higher recurrence rate Radiation exposure

REFERENCES

1. Macey MR, Owen RC, Ross SS, Coward RM. Best practice in the diagnosis and treatment of varicocele in children and adolescents. *Ther Adv Urol.* 2018 Jun 22;10(9):273-282. doi: 10.1177/1756287218783900.
2. Glassberg KI, Korets R. Update on the management of adolescent varicocele. *F1000 Med Rep.* 2010 Apr 12;2:25. doi: 10.3410/M2-25.
3. Cho PS, Yu RN, Paltiel HJ, Migliozi MA, Li X, Venna A, Diamond DA. Clinical outcome of pediatric and young adult subclinical varicoceles: a single-institution experience. *Asian J Androl.* 2021 Nov-Dec;23(6):611-615. doi: 10.4103/aja.aja_22_21.
4. Diamond DA, Zurakowski D, Bauer SB, Borer JG, Peters CA, Cilento BG Jr, Paltiel HJ, Rosoklija I, Retik AB. Relationship of varicocele grade and testicular hypotrophy to semen parameters in adolescents. *J Urol.* 2007 Oct;178(4 Pt 2):1584-8. doi: 10.1016/j.juro.2007.03.169. Epub 2007 Aug 16.
5. Zhou T, Zhang W, Chen Q, Li L, Cao H, Xu CL, Chen GH, Sun YH. Effect of varicolectomy on testis volume and semen parameters in adolescents: a meta-analysis. *Asian J Androl.* 2015 Nov-Dec;17(6):1012-6. doi: 10.4103/1008-682X.148075.
6. Moursy EE, Eldahshoury MZ, Hussein MM, Mourad MZ, Badawy AA. Dilemma of adolescent varicocele: long-term outcome in patients managed surgically and in patients managed expectantly. *J Pediatr Urol.* 2013 Dec;9(6 Pt B):1018-22. doi: 10.1016/j.jpuro.2013.01.017. Epub 2013 Mar 20.
7. Radmayr C, Bogart G, Burgu B, Castagnetti MS, Dogan HS, O’Kelly F, Quaedackers J, Rawas-hdeh YFH, Silay MS. EAU Guideline on Pediatric Urology- <https://uroweb.org/guidelines/paediatric-urology>, Limited Update March, 2023.
8. Lourdaux PJ, Vaganée D, Leysen C, De Wachter S, De Win G. Evolution of testicular as-

- ymmetry during puberty in adolescents without and with a left varicocele. *BJU Int.* 2023 Mar;131(3):348-356. doi: 10.1111/bju.15914. Epub 2022 Nov 1.
9. Zundel S, Szavay P, Stanasel I. Management of adolescent varicocele. *Semin Pediatr Surg.* 2021 Aug;30(4):151084. doi: 10.1016/j.sempedsurg.2021.151084. Epub 2021 Jul 14.
 10. Vaganée D, Daems F, Aerts W, Dewaide R, van den Keybus T, De Baets K, De Wachter S, De Win G. Testicular asymmetry in healthy adolescent boys. *BJU Int.* 2018 Oct;122(4):654-666. doi: 10.1111/bju.14174. Epub 2018 Mar 23.
 11. Zampieri N. Hormonal evaluation in adolescents with varicocele. *J Pediatr Urol.* 2021 Feb;17(1):49.e1-49.e5. doi: 10.1016/j.jpuro.2020.11.024. Epub 2020 Nov 21.
 12. Zampieri N. Varicocele in adolescents: new findings and certainties among volume, hormones and semen analysis. Open perspectives. *Am J Clin Exp Urol.* 2021 Jun 15;9(3):239-241.
 13. Bogaert GA. Adolescent varicocele: limited indications for treatment during puberty and adolescence. *Transl Androl Urol.* 2014 Dec;3(4):398-401. doi: 10.3978/j.issn.2223-4683.2014.12.10.
 14. Paduch DA, Niedzielski J. Repair versus observation in adolescent varicocele: a prospective study. *J Urol.* 1997 Sep;158(3 Pt 2):1128-32. doi: 10.1097/00005392-199709000-00111.
 15. Cannarella R, Calogero AE, Condorelli RA, Giaccone F, Aversa A, La Vignera S. Management and Treatment of Varicocele in Children and Adolescents: An Endocrinologic Perspective. *J Clin Med.* 2019 Sep 8;8(9):1410. doi: 10.3390/jcm8091410.
 16. Nork JJ, Berger JH, Crain DS, Christman MS. Youth varicocele and varicocele treatment: a meta-analysis of semen outcomes. *Fertil Steril.* 2014 Aug;102(2):381-387.e6. doi: 10.1016/j.fertnstert.2014.04.049. Epub 2014 Jun 4.
 17. Chung JM, Lee SD. Current Issues in Adolescent Varicocele: Pediatric Urological Perspectives. *World J Mens Health.* 2018 May;36(2):123-131. doi: 10.5534/wjmh.170053. Epub 2018 Mar 22.
 18. Silay MS, Hoen L, Quadackaers J, Undre S, Bogaert G, Dogan HS, Kocvara R, Nijman RJM, Radmayr C, Tekgul S, Stein R. Treatment of Varicocele in Children and Adolescents: A Systematic Review and Meta-analysis from the European Association of Urology/European Society for Paediatric Urology Guidelines Panel. *Eur Urol.* 2019 Mar;75(3):448-461. doi: 10.1016/j.eururo.2018.09.042.
 19. Elder J. Does the Evidence Support Adolescent Varicocelectomy? *Eur Urol.* 2019 Mar;75(3):462-463. doi: 10.1016/j.eururo.2018.10.059. Epub 2018 Nov 22.
 20. Cayan S, Kadioglu A, Orhan I, Kandirali E, Tefekli A, Tellaloglu S. The effect of microsurgical varicocelectomy on serum follicle stimulating hormone, testosterone and free testosterone levels in infertile men with varicocele. *BJU Int.* 1999 Dec;84(9):1046-9. doi: 10.1046/j.1464-410x.1999.00353.x.
 21. Takeyama M, Honjoh M, Kodama M, Sakaguchi H, Koh E, Kondoh N, Fujioka H, Nakamura M, Namiki M, Okuyama A, et al. Testicular steroids in spermatid and peripheral veins after single injection of hCG in patients with varicocele. *Arch Androl.* 1990;24(2):207-13. doi: 10.3109/01485019008986881.
 22. Fujisawa M, Dobashi M, Yamasaki T, Kanzaki M, Okada H, Arakawa S, Kamidono S. Significance of serum inhibin B concentration for evaluating improvement in spermatogenesis after varicocelectomy. *Hum Reprod.* 2001 Sep;16(9):1945-9. doi: 10.1093/humrep/16.9.1945.
 23. Patil N, Javali T. Varicocelectomy in adolescents - Does it safeguard future fertility? A single centre experience. *J Pediatr Urol.* 2022 Feb;18(1):5.e1-5.e10. doi: 10.1016/j.jpuro.2021.11.020. Epub 2021 Dec 5.
 24. Laven JS, Haans LC, Mali WP, te Velde ER, Wensing CJ, Eimers JM. Effects of varicocele treatment in adolescents: a randomized study. *Fertil Steril.* 1992 Oct;58(4):756-62. doi: 10.1016/s0015-0282(16)55324-2.

25. Van Batavia JP, Fast AM, Nees SN, Mercado MA, Gaselberti A, Glassberg KI. Incidence, significance and natural history of persistent retrograde venous flow after varicocelectomy in children and adolescents: correlation with catch-up growth. *J Urol.* 2013 Aug;190(2):689-95. doi: 10.1016/j.juro.2013.02.3195.
26. Li F, Chiba K, Yamaguchi K, Okada K, Matsushita K, Ando M, Yue H, Fujisawa M. Effect of varicocelectomy on testicular volume in children and adolescents: a meta-analysis. *Urology.* 2012 Jun;79(6):1340-5. doi: 10.1016/j.urology.2012.02.022. Epub 2012 Apr 17.
27. Okuyama A, Nakamura M, Namiki M, Takeyama M, Utsunomiya M, Fujioka H, Itatani H, Matsuda M, Matsumoto K, Sonoda T. Surgical repair of varicocele at puberty: preventive treatment for fertility improvement. *J Urol.* 1988 Mar;139(3):562-4. doi: 10.1016/s0022-5347(17)42525-0.
28. Chiba K, Ramasamy R, Lamb DJ, Lipshultz LI. The varicocele: diagnostic dilemmas, therapeutic challenges and future perspectives. *Asian J Androl.* 2016 Mar-Apr;18(2):276-81. doi: 10.4103/1008-682X.167724.
29. Franco A, Proietti F, Palombi V, Savarese G, Guidotti M, Leonardo C, Ferro F, Manna C, Franco G. Varicocele: To Treat or Not to Treat? *J Clin Med.* 2023 Jun 15;12(12):4062. doi: 10.3390/jcm12124062.
30. Bogaert G, Orye C, De Win G. Pubertal screening and treatment for varicocele do not improve chance of paternity as adult. *J Urol.* 2013 Jun;189(6):2298-303. doi: 10.1016/j.juro.2012.12.030. Epub 2012 Dec 20.
31. Chu DI, Zderic SA, Shukla AR, Srinivasan AK, Tasian GE, Weiss DA, Long CJ, Canning DA, Kolon TF. The natural history of semen parameters in untreated asymptomatic adolescent varicocele patients: A retrospective cohort study. *J Pediatr Urol.* 2017 Feb;13(1):77.e1-77.e5. doi: 10.1016/j.jpuro.2016.09.008. Epub 2016 Oct 24.
32. Sarah C. Krzastek, Connor Rotterman, Ryan P. Smith, and Jason R. Kovac. Macroscopic Surgical Techniques for Varicocele Repair. Chapter 16, pp: 201-208, Varicocele and Male Infertility Eds: Sandro C. Esteves, Chak-Lam Cho, Ahmad Majzoub, Ashok Agarwal, Springer Nature Switzerland AG 2019
33. Palomo A. Radical cure of varicocele by a new technique; preliminary report. *J Urol.* 1949 Mar;61(3):604-7. doi: 10.1016/S0022-5347(17)69113-4.
34. Paick S, Choi WS. Varicocele and Testicular Pain: A Review. *World J Mens Health.* 2019 Jan;37(1):4-11. doi: 10.5534/wjmh.170010. Epub 2018 May 16.
35. Tsai CS, Lin FY, Chen YH, Yang TL, Wang HJ, Huang GS, Lin CY, Tsai YT, Lin SJ, Li CY. Cilostazol attenuates MCP-1 and MMP-9 expression in vivo in LPS-administrated balloon-injured rabbit aorta and in vitro in LPS-treated monocytic THP-1 cells. *J Cell Biochem.* 2008 Jan 1;103(1):54-66. doi: 10.1002/jcb.21388.
36. Lundy SD, Sabanegh ES Jr. Varicocele management for infertility and pain: A systematic review. *Arab J Urol.* 2017 Dec 14;16(1):157-170. doi: 10.1016/j.aju.2017.11.003.
37. Parrilli A, Roberti A, Escolino M, Esposito C. Surgical approaches for varicocele in pediatric patient. *Transl Pediatr.* 2016 Oct;5(4):227-232. doi: 10.21037/tp.2016.09.11.
38. Misseri R, Gershbein AB, Horowitz M, Glassberg KI. The adolescent varicocele. II: the incidence of hydrocele and delayed recurrent varicocele after varicocelectomy in a long-term follow-up. *BJU Int.* 2001 Apr;87(6):494-8. doi: 10.1046/j.1464-410x.2001.00110.x.
39. Ivanissevich O. Left varicocele due to reflux; experience with 4,470 operative cases in forty-two years. *J Int Coll Surg.* 1960 Dec;34:742-55.
40. Hopps CV, Lemer ML, Schlegel PN, Goldstein M. Intraoperative varicocele anatomy: a microscopic study of the inguinal versus subinguinal approach. *J Urol.* 2003 Dec;170(6 Pt 1):2366-70. doi: 10.1097/01.ju.0000097400.67715.f8.
41. Liu WL, Chang JM, Chong IW, Hung YL, Chen YH, Huang WT, Kuo HF, Hsieh CC, Liu PL.

- Curcumin Inhibits LIN-28A through the Activation of miRNA-98 in the Lung Cancer Cell Line A549. *Molecules*. 2017 Jun 3;22(6):929. doi: 10.3390/molecules22060929.
42. Çayan S, Şahin S, Akbay E. Paternity Rates and Time to Conception in Adolescents with Varicocele Undergoing Microsurgical Varicocele Repair vs Observation Only: A Single Institution Experience with 408 Patients. *J Urol*. 2017 Jul;198(1):195-201. doi: 10.1016/j.juro.2017.01.066. Epub 2017 Jan 31.
 43. Owen RC, McCormick BJ, Figler BD, Coward RM. A review of varicocele repair for pain. *Transl Androl Urol*. 2017 May;6(Suppl 1):S20-S29. doi: 10.21037/tau.2017.03.36.
 44. Zampieri N, Cervellione RM. Varicocele in adolescents: a 6-year longitudinal and followup observational study. *J Urol*. 2008 Oct;180(4 Suppl):1653-6; discussion 1656. doi: 10.1016/j.juro.2008.03.114. Epub 2008 Aug 20.
 45. Harel M, Herbst KW, Nelson E. Practice patterns in the surgical approach for adolescent varicolectomy. *Springerplus*. 2015 Dec 14;4:772. doi: 10.1186/s40064-015-1573-7.
 46. Podkamenev VV, Stalmakhovich VN, Urkov PS, Solovjev AA, Iljin VP. Laparoscopic surgery for pediatric varicoceles: Randomized controlled trial. *J Pediatr Surg*. 2002 May;37(5):727-9. doi: 10.1053/jpsu.2002.32264.
 47. Rotker K, Sigman M. Recurrent varicocele. *Asian J Androl*. 2016 Mar-Apr;18(2):229-33. doi: 10.4103/1008-682X.171578.
 48. Ding H, Tian J, Du W, Zhang L, Wang H, Wang Z. Open non-microsurgical, laparoscopic or open microsurgical varicolectomy for male infertility: a meta-analysis of randomized controlled trials. *BJU Int*. 2012 Nov;110(10):1536-42. doi: 10.1111/j.1464-410X.2012.11093.x. Epub 2012 May 29.
 49. Park JH, Pak K, Park NC, Park HJ. How Can We Predict a Successful Outcome after Varicolectomy in Painful Varicocele Patients? An Updated Meta-Analysis. *World J Mens Health*. 2021 Oct;39(4):645-653. doi: 10.5534/wjmh.190112. Epub 2019 Dec 3.
 50. Richter F, Stock JA, LaSalle M, Sadeghi-Nejad H, Hanna MK. Management of prepubertal varicoceles-results of a questionnaire study among pediatric urologists and urologists with infertility training. *Urology*. 2001 Jul;58(1):98-102. doi: 10.1016/s0090-4295(01)01118-9.
 51. Pastuszak AW, Kumar V, Shah A, Roth DR. Diagnostic and management approaches to pediatric and adolescent varicocele: a survey of pediatric urologists. *Urology*. 2014 Aug;84(2):450-5. doi: 10.1016/j.urology.2014.04.022. Epub 2014 Jun 11.
 52. Lee TH, Jung JH, Hong YK. Diagnosis and Management of Pediatric and Adolescent Varicocele: A Survey of Pediatric Urologists in Korea. *Chonnam Med J*. 2016 Sep;52(3):207-11. doi: 10.4068/cmj.2016.52.3.207. Epub 2016 Sep 23.
 53. Tauber R, Johnsen N. Die antegrade skrotale Verödung zur Behandlung der Testisvarikozele. Technik und Spätergebnisse [Antegrade scrotal sclerotherapy for treatment of testicular varicocele. Technique and late results]. *Urologe A*. 1993 Jul;32(4):320-6. German.
 54. Cassidy D, Jarvi K, Grober E, Lo K. Varicocele surgery or embolization: Which is better? *Can Urol Assoc J*. 2012 Aug;6(4):266-8. doi: 10.5489/cuaj.11064.
 55. Zampieri N, Chironi C, Sulpasso M. Treatment of varicocele with transfemoral retrograde sclero-embolization in pediatric patients under local anesthesia. *Minerva Pediatr*. 2015 Jun;67(3):227-9. Epub 2014 Jul 11.
 56. Hawkins CM, Racadio JM, McKinney DN, Racadio JM, Vu DN. Varicocele retrograde embolization with boiling contrast medium and gelatin sponges in adolescent subjects: a clinically effective therapeutic alternative. *J Vasc Interv Radiol*. 2012 Feb;23(2):206-10. doi: 10.1016/j.jvir.2011.10.021. Epub 2011 Dec 15.
 57. Malekzadeh S, Fraga-Silva RA, Morère PH, Sorega A, Produit S, Stergiopoulos N, Constantin C. Varicocele percutaneous embolization outcomes in a pediatric group: 7-year retrospective study. *Int Urol Nephrol*. 2016 Sep;48(9):1395-9. doi: 10.1007/s11255-016-1340-x. Epub 2016

- Jun 30.
58. Wickham A, Vu D, El-Arabi A, Gatti JM. Adolescent Varicocelectomy: Success at What Cost? Clinical Outcome and Cost Comparison of Surgical Ligation and Percutaneous Embolization. *J Laparoendosc Adv Surg Tech A*. 2021 Aug;31(8):942-946. doi: 10.1089/lap.2020.0836. Epub 2021 Jul 9.
 59. Tauber R, Pfeiffer D. Surgical atlas varicocele: Antegrade scrotal sclerotherapy. *BJU Int*. 2006 Dec;98(6):1333-44. doi: 10.1111/j.1464-410X.2006.06579.x.
 60. Keene DJB, Cervellione RM. Antegrade sclerotherapy in adolescent varicocele patients. *J Pediatr Urol*. 2017 Jun;13(3):305.e1-305.e6. doi: 10.1016/j.jpuro.2016.12.018. Epub 2017 Jan 29.
 61. Mazzoni G, Spagnoli A, Lucchetti MC, Villa M, Capitanucci ML, Ferro F. Adolescent varicocele: Tauber antegrade sclerotherapy versus Palomo repair. *J Urol*. 2001 Oct;166(4):1462-4. doi: 10.1016/s0022-5347(05)65810-7.
 62. Fabiani A, Pavia MP, Stramucci S, Antezza A, De Stefano V, Castellani D. Do sclero-embolization procedures have advantages over surgical ligation in treating varicocele in children, adolescents and adults? Results from a systematic review and meta-analysis. *Andrologia*. 2022 Sep;54(8):e14510. doi: 10.1111/and.14510. Epub 2022 Jun 24.
 63. Halpern J, Mittal S, Pereira K, Bhatia S, Ramasamy R. Percutaneous embolization of varicocele: technique, indications, relative contraindications, and complications. *Asian J Androl*. 2016 Mar-Apr;18(2):234-8. doi: 10.4103/1008-682X.169985.
 64. Çayan S, Bozlu M, Akbay E. Update on the novel management and future paternity situation in adolescents with varicocele. *Turk J Urol*. 2017 Sep;43(3):241-246. doi: 10.5152/tud.2017.01033. Epub 2017 Aug 1.