## GRADE Table 1. Effectiveness of one dose varicella vaccination in immunocompetent children (9 months to 12 years of age) in preventing all grades of severity of varicella (evidence available for within the first 10 years after vaccination)

**Population**: Immunocompetent children (9 month to 12 years of age)

**Intervention:** One-dose varicella vaccination **Comparison:** Placebo/ No vaccination

**Outcome**: All grades of severity of varicella disease

What is the scientific evidence of the effectiveness of one dose of varicella vaccination (versus placebo/no vaccination) in preventing all grades of severity of varicella in immunocompetent children (9 months to 12 years of age)?

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			Rating	Adjustment to rating
Quality Assessment	No. of studies/starting rating		40/ Observational <sup>1</sup>	2
	Factors decreasing confidence	Limitation in study design	None serious	0
		Inconsistency	None serious	0
		Indirectness	None serious	0
		Imprecision	None serious	0
		Publication bias	None serious	0
	Factors increasing confidence	Large effect <sup>2</sup>	Applicable	+2
		Dose-response	Not applicable	0
		Antagonistic bias and confounding	Not applicable	0
	Final numerical rating of quality of evidence			4
10	Statement on quality of evidence			We are very confident that the true effect lies close to that of the estimate of effect on health outcome
Summary of Findings	Conclusion			A single dose of varicella vaccination is effective to protect children of 9 months to 12 years against all grades of severity of varicella disease. Single dose varicella VE against all grades of disease severity ranged from 20 – 100%, with an approximate mean VE of 80% against all grades of disease severity, irrespective of vaccine type.

## Reference List<sup>1-45</sup>

 $<sup>^1</sup>$  Two systematic reviews (Seward et al.; Bayer et al.) and a syst. rev. done by WHO of the current literature (through October 2013) identified 40 observational studies. Single dose varicella VE against all grades of disease severity ranged from 20 – 100%, with an approximate mean VE of 80% against all grades of disease severity, irrespective of vaccine type. Only one study demonstrated vaccine effectiveness against all varicella to be 20% (95%CI 0%-40%).

<sup>&</sup>lt;sup>2</sup> Upgraded by two levels as strong evidence from observational studies of a vaccine effectiveness of 80% or higher with no major residual confounders. In addition to effectiveness on an individual level, decline in incidence in all age groups over time, not only age-group targeted by vaccination program, suggests induction of community protection (Marin et al 2008, Marin et al 2011, Lopez et al 2011, Guris et al 2008).

- 1. Seward JF, Marin M, Vazquez M. Varicella vaccine effectiveness in the US vaccination program: a review. J Infect Dis 2008;197 Suppl 2:S82-S89.
- 2. Outbreak of varicella among vaccinated children--Michigan, 2003 27. MMWR Morb Mortal Wkly Rep 2004;53(18):389-392.
- 3. Varicella outbreak among vaccinated children--Nebraska, 2004 22. MMWR Morb Mortal Wkly Rep 2006;55(27):749-752.
- 4. Arnedo-Pena A, Puig-Barbera J, Aznar-Orenga MA et al. Varicella vaccine effectiveness during an outbreak in a partially vaccinated population in Spain 4. Pediatr Infect Dis J 2006;25(9):774-778.
- 5. Buchholz U, Moolenaar R, Peterson C, Mascola L. Varicella outbreaks after vaccine licensure: should they make you chicken? 1. Pediatrics 1999;104(3 Pt 1):561-563.
- 6. Cenoz MG, Martinez-Artola V, Guevara M, Ezpeleta C, Barricarte A, Castilla J. Effectiveness of one and two doses of varicella vaccine in preventing laboratory-confirmed cases in children in Navarre, Spain 1. Hum Vaccin Immunother 2013;9(5).
- 7. Clements DA, Moreira SP, Coplan PM, Bland CL, Walter EB. Postlicensure study of varicella vaccine effectiveness in a day-care setting. Pediatr Infect Dis J 1999;18(12):1047-1050.
- 8. Dworkin MS, Jennings CE, Roth-Thomas J, Lang JE, Stukenberg C, Lumpkin JR. An Outbreak of Varicella among children attending preschool and elementary school in Illinois. Clin Infect Dis 2002;35(1):102-104.
- 9. Fu C, Wang M, Liang J, Xu J, Wang C, Bialek S. The effectiveness of varicella vaccine in China 1. Pediatr Infect Dis J 2010;29(8):690-693.
- 10. Galil K, Lee B, Strine T et al. Outbreak of varicella at a day-care center despite vaccination. N Engl J Med 2002;347(24):1909-1915.
- 11. Galil K, Fair E, Mountcastle N, Britz P, Seward J. Younger age at vaccination may increase risk of varicella vaccine failure 8. J Infect Dis 2002;186(1):102-105.
- 12. Gould PL, Leung J, Scott C et al. An outbreak of varicella in elementary school children with two-dose varicella vaccine recipients--Arkansas, 2006. Pediatr Infect Dis J 2009;28(8):678-681.
- 13. Haddad MB, Hill MB, Pavia AT et al. Vaccine effectiveness during a varicella outbreak among schoolchildren: Utah, 2002-2003 1. Pediatrics 2005;115(6):1488-1493.
- 14. Hohle M, Siedler A, Bader HM, Ludwig M, Heininger U, Von KR. Assessment of varicella vaccine effectiveness in Germany: a time-series approach 1. Epidemiol Infect 2011;139(11):1710-1719.
- 15. Huang WC, Huang LM, Chang IS, Tsai FY, Chang LY. Varicella breakthrough infection and vaccine effectiveness in Taiwan 4. Vaccine 2011;29(15):2756-2760.
- 16. Izurieta HS, Strebel PM, Blake PA. Postlicensure effectiveness of varicella vaccine during an outbreak in a child care center 5. JAMA 1997;278(18):1495-1499.

- 17. Kilic A, Unuvar E, Yilmaz C, Yildiz I, Oguz F, Sidal M. The effectiveness of varicella vaccination during an outbreak in a children's day-care center 2. Vaccine 2008;26(27-28):3371-3372.
- 18. Lai CC, Chen SC, Jiang DD. An outbreak of varicella among schoolchildren in Taipei. BMC Public Health 2011;11:226.
- 19. Lee BR, Feaver SL, Miller CA, Hedberg CW, Ehresmann KR. An elementary school outbreak of varicella attributed to vaccine failure: policy implications 9. J Infect Dis 2004;190(3):477-483.
- 20. Lee LE, Ho H, Lorber E, Fratto J, Perkins S, Cieslak PR. Vaccine-era varicella epidemiology and vaccine effectiveness in a public elementary school population, 2002-2007 52. Pediatrics 2008;121(6):e1548-e1554.
- 21. Liese JG, Cohen C, Rack A et al. The effectiveness of varicella vaccination in children in Germany: a case-control study 2. Pediatr Infect Dis J 2013;32(9):998-1004.
- 22. Lopez AS, Guris D, Zimmerman L et al. One dose of varicella vaccine does not prevent school outbreaks: is it time for a second dose? 5. Pediatrics 2006;117(6):e1070-e1077.
- 23. Lu L, Suo L, Li J et al. A varicella outbreak in a school with high one-dose vaccination coverage, Beijing, China. Vaccine 2012;30(34):5094-5098.
- 24. Mahamud A, Wiseman R, Grytdal S et al. Challenges in confirming a varicella outbreak in the two-dose vaccine era 1. Vaccine 2012;30(48):6935-6939.
- 25. Marin M, Nguyen HQ, Keen J et al. Importance of catch-up vaccination: experience from a varicella outbreak, Maine, 2002-2003 8. Pediatrics 2005;115(4):900-905.
- 26. Miron D, Lavi I, Kitov R, Hendler A. Vaccine effectiveness and severity of varicella among previously vaccinated children during outbreaks in day-care centers with low vaccination coverage. Pediatr Infect Dis J 2005;24(3):233-236.
- 27. Nguyen MD, Perella D, Watson B, Marin M, Renwick M, Spain CV. Incremental effectiveness of second dose varicella vaccination for outbreak control at an elementary school in Philadelphia, pennsylvania, 2006 2. Pediatr Infect Dis J 2010;29(8):685-689.
- 28. Parker AA, Reynolds MA, Leung J et al. Challenges to implementing second-dose varicella vaccination during an outbreak in the absence of a routine 2-dose vaccination requirement-Maine, 2006 2. J Infect Dis 2008;197 Suppl 2:S101-S107.
- 29. Passwell JH, Hemo B, Levi Y, Ramon R, Friedman N, Lerner-Geva L. Use of a computerized database to study the effectiveness of an attenuated varicella vaccine 2. Pediatr Infect Dis J 2004;23(3):221-226.
- 30. Seward JF, Zhang JX, Maupin TJ, Mascola L, Jumaan AO. Contagiousness of varicella in vaccinated cases: a household contact study. JAMA 2004;292(6):704-708.
- 31. Shapiro ED, Vazquez M, Esposito D et al. Effectiveness of 2 doses of varicella vaccine in children. J Infect Dis 2011;203(3):312-315.

- 32. Sheffer R, Segal D, Rahamani S et al. Effectiveness of the Oka/GSK attenuated varicella vaccine for the prevention of chickenpox in clinical practice in Israel 1. Pediatr Infect Dis J 2005;24(5):434-437.
- 33. Spackova M, Wiese-Posselt M, Dehnert M, Matysiak-Klose D, Heininger U, Siedler A. Comparative varicella vaccine effectiveness during outbreaks in day-care centres 2. Vaccine 2010;28(3):686-691.
- 34. Tafuri S, Martinelli D, Prato R, Germinario C. Vaccine effectiveness evaluation during a varicella outbreak among children of primary schools and day-care centers in a region which adopted UMV 1. Hum Vaccin Immunother 2013;9(1):184-188.
- 35. Tafuri S, Martinelli D, De PM, Germinario C, Prato R. Report of varicella outbreak in a low vaccination coverage group of otherwise healthy children in Italy: the role of breakthrough and the need of a second dose of vaccine. Vaccine 2010;28(6):1594-1597.
- 36. Tugwell BD, Lee LE, Gillette H, Lorber EM, Hedberg K, Cieslak PR. Chickenpox outbreak in a highly vaccinated school population 1. Pediatrics 2004;113(3 Pt 1):455-459.
- 37. Vally H, Dowse GK, Eastwood K, Cameron S. An outbreak of chickenpox at a child care centre in Western Australia. Costs to the community and implications for vaccination policy 2. Aust N Z J Public Health 2007;31(2):113-119.
- 38. Vazquez M, LaRussa PS, Gershon AA, Steinberg SP, Freudigman K, Shapiro ED. The effectiveness of the varicella vaccine in clinical practice. N Engl J Med 2001;344(13):955-960.
- 39. Vazquez M, LaRussa PS, Gershon AA et al. Effectiveness over time of varicella vaccine 2. JAMA 2004;291(7):851-855.
- 40. Wang Z, Yang H, Li K et al. Single-dose varicella vaccine effectiveness in school settings in China 6. Vaccine 2013;31(37):3834-3838.
- 41. Bayer O, Heininger U, Heiligensetzer C, Von KR. Metaanalysis of vaccine effectiveness in varicella outbreaks 1. Vaccine 2007;25(37-38):6655-6660.
- 42. Marin M, Meissner HC, Seward JF. Varicella prevention in the United States: a review of successes and challenges 3. Pediatrics 2008;122(3):e744-e751.
- 43. Marin M, Zhang JX, Seward JF. Near elimination of varicella deaths in the US after implementation of the vaccination program 1. Pediatrics 2011;128(2):214-220.
- 44. Lopez AS, Zhang J, Brown C, Bialek S. Varicella-related hospitalizations in the United States, 2000-2006: the 1-dose varicella vaccination era 5. Pediatrics 2011;127(2):238-245.
- 45. Guris D, Jumaan AO, Mascola L et al. Changing varicella epidemiology in active surveillance sites--United States, 1995-2005 1. J Infect Dis 2008;197 Suppl 2:S71-S75.