

**Table V: Sequential administration IPV-OPV****Population :** Immunocompetent individuals**Intervention:** Sequential administration of IPV followed by OPV**Comparison:** No vaccination**Outcome :** Immunogenicity/ Cases of poliomyelitis/ VAPP

<b>PICO Question:</b> What is the quality of scientific evidence that sequential immunization schedules starting with $\geq 2$ doses of IPV followed by $\geq 2$ doses of OPV induce protective immune responses to all two poliovirus serotypes in $\geq 90\%$ of vaccines, (i.e. responses comparable to those induced by the same number of doses of either OPV or IPV alone				
		Rating	Adjustment to rating	
<b>Quality Assessment</b>	No of studies/starting rating		2 RCT/ 6 observational <sup>1</sup>	4
	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 detected	0
		Factors increasing confidence	Strength of association/ large effect	Not applicable
	Dose-response		Not applicable	0
	Antagonistic /mitigated bias and confounding		Not applicable	0
	<b>Final numerical rating of quality of evidence</b>			<b>4</b>
<b>Summary of Findings</b>	<b>Statement on quality of evidence</b>		Evidence supports a high degree of confidence that the true effect lies close to that of the estimate of effect on health outcome.	
	<b>Conclusion</b>		High scientific evidence that a sequential schedule of IPV and OPV protects against clinical poliomyelitis.	

<sup>1</sup>Modlin JF et al (1997) showed in a randomized controlled study of 510 infants, that for each of the 3 IPV-OPV experimental sequential schedules, the first OPV dose significantly enhanced seroconversion rates and geometric mean micro-neutralization antibody titers (GMT). 3 months after the final dose, 96%-99%, 99%-100%, and 81%-100% of infants had antibodies to poliovirus types 1, 2, and 3, respectively. Subjects with  $\geq 2$  prior OPV doses were significantly less likely than those with none or one prior OPV dose to excrete virus in feces after an OPV challenge. Faden H et al (1990) showed that incorporation of at least one dose of IPV at the start of the immunization schedule tends to increase systemic as well as local antibody production. Faden H et al (1993) showed that as compared to OPV-OPV-OPV, eIPV-eIPV-eIPV, eIPV-OPV-OPV, and eIPV-eIPV-OPV those receiving the eIPV-eIPV-OPV schedule maintained the highest antibody levels throughout the observation period, including the response to OPV challenge at 5 years of age. Swartz TA et al (1998) assessing the effectiveness of an intercalated IPV-OPV vaccine programme in Israel concluded that the programme offered high individual protection throughout the first 5 years of life. von Magnus H et al (1984) reported that in Denmark with a sequential 3-dose IPV/OPV immunization programme since 1968,  $\geq 95\%$  of the population had antibodies to poliovirus, and the GMT of serum antibodies  $\geq 10$  IU for all three types. Lu CY et al (2001) showed that protective antibodies were present in all infants (6 months), 2 months after the second IPV dose. Antibody titers were augmented at the age of 19 months, 1 month after the booster dose of OPV. In the US no VAPP case occurred with the IPV-OPV schedule (Alexander et al. 2004). Seroconversion rates against polioviruses types 1 and 3 were non-inferior in sequential schedules containing IPV and bOPV, compared with an all-IPV schedule, and proportions of infants with protective antibodies were high (O’Ryan et al 2015).

## References

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