Varicella More than just a rash

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Often considered a benign childhood illness, varicella (chickenpox) can be associated with serious and lifethreatening complications. GPs have an important role in providing varicella vaccinations and in identifying patients with varicella infection or exposure, especially high-risk individuals who require counselling, prophylaxis and possible hospital referral.

aricella (also known as chickenpox) is a highly contagious viral illness caused by primary infection with varicella zoster virus (VZV). It is characterised by general malaise and fever followed by the eruption of a vesicular rash over the next three to five days.¹⁻³ Reactivation of VZV causes herpes zoster (shingles), a disease most prevalent in people aged over 60 years.⁴

Chickenpox is commonly seen in general practice among unvaccinated individuals. Often considered a benign childhood illness, it can be associated with serious and life-threatening complications, including pneumonia and disseminated disease.

This article focuses on chickenpox in childhood. It summarises the role of varicella vaccination and management of individuals at high risk of serious complications, including postexposure prophylaxis and antiviral treatment of chickenpox.

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Epidemiology

Chickenpox is common in children aged under 15 years, with more than 90% of the unvaccinated population in temperate climates developing clinical infection or subclinical seroconversion by adolescence.^{5,6} Before the introduction of universal vaccination, there were about 240,000 cases and 1500 hospitalisations each year from varicella in Australia.⁷

After the introduction of a single-dose varicella vaccine under the National Immunisation Program (NIP) in 2005, hospitalisations for chickenpox declined rapidly in all age groups under 40 years. ^{2,7} The greatest reduction (84%) was in children aged 18 to 59 months. In 2016, there were 395 varicella-coded hospitalisations, of which 101 (26%) were in children aged under 15 years.¹

Between 1997 and 2016, 132 deaths were recorded from chickenpox in Australia. Among these, 15 deaths were in children aged under 15 years, only two of which occurred after vaccination commenced.¹ The rates of varicella-related hospitalisations and deaths in Australia before and after the introduction of varicella vaccine are shown in Figures 1a and b.¹

Pathogenesis

VZV is a double-stranded DNA virus and member of the herpes virus family.⁸ VZV infection causes two distinct clinical conditions:

- Primary infection with VZV causes varicella (chickenpox), with subsequent lifetime latency of VZV in the dorsal root ganglion.⁹
- Reactivation of latent VZV typically results in a discrete rash in a dermatomal distribution, known as herpes zoster (shingles).^{7,8}



Figures 1a and b. Rate of varicella-related hospitalisations and deaths in Australia before and after the introduction of varicella vaccine.

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VZV can be transmitted via respiratory droplets before the development of the vesicular rash or via direct contact with vesicular fluid. The incubation period ranges from 10 to 21 days (most commonly 14 to 16 days). However, the incubation period may be prolonged up to 28 days in patients who receive zoster immunoglobulin.¹⁰ Individuals are infectious from one to two days before onset of the rash until all lesions have crusted over. Figure 2 shows a typical varicella rash with lesions at different stages of development.^{5,8}

Complications

Chickenpox is typically a mild and selflimiting illness in healthy children. Complications occur in about 1% of cases and are more common in neonates, adults and individuals who are immunocompromised by another medical condition or its treatment.^{5,8}

Complications of chickenpox include secondary bacterial infections (5%), pneumonia (1%), thrombocytopenia and neurological problems (1 to 3.5 in every 100,000 children aged under 14 years).¹¹ Immunocompromised individuals are at higher risk of encephalomyelitis, cerebellar ataxia, hepatitis, arthritis, haemorrhagic nephritis, myocarditis and otitis media.^{7,11} Secondary bacterial infections, particularly invasive group A streptococcal or staphylococcal infections (e.g. cellulitis, myositis, necrotising fasciitis and toxic shock syndrome), are the most common cause of hospital admission in children.^{11,12}

Management

The mainstay of treatment of chickenpox is supportive care, including hydration, antipyretics and pruritis management, as well as monitoring for complications.^{3,10} Calamine lotion and cool compresses can alleviate itching.¹⁰ Salicylates (aspirin) should be avoided because of the risk of Reye syndrome, an extremely rare acute encephalopathy with associated hepatic dysfunction. Reye syndrome is characterised by vomiting and confusion and may rapidly evolve to seizures, coma and death.¹²

Antiviral therapies licensed for chickenpox in children and adolescents include aciclovir and valaciclovir. Famciclovir is not used in children.¹³ Chickenpox in pregnant women, neonates, immunocompromised patients or patients with severe or complicated disease should prompt urgent referral for specialist and hospital care for consideration of antiviral therapy.14,15

Chickenpox is currently notifiable in all Australian states and territories apart from NSW.¹

Prevention

The current recommended VZV vaccination schedule, adapted from the Australian Immunisation Handbook, is shown in the Table.14,16,17

Varicella vaccine

Varicella vaccine is a live attenuated vaccine that has been included in the Australian NIP since November 2005.7,8 It is given routinely as a single dose in combination with measles, mumps and rubella vaccines (MMRV) at 18 months of age.7,14 A second dose of varicella vaccine, not currently funded by the NIP, is recommended to improve protection and minimise the chance of breakthrough disease in children under 14 years of age.14

Single-dose varicella vaccine was shown to be 81% effective in preventing any chickenpox and 98% effective in preventing

Figure 2. Varicella rash showing lesions at different stages of development, with many lesions crusting. Image courtesy of The Royal Children's Hospital Melbourne (www.rch.org.au/ clinicalguide/guideline_ index/Chickenpox_ varicella/).



moderate to severe disease.18 Effectiveness in preventing any chickenpox rises to 92% after two doses of varicella vaccine. Varicella vaccine also reduces the risk of herpes zoster among vaccinated children.¹⁷⁻¹⁹ However, the rate of primary vaccine failure ranges from 9% to 14% after one dose, and the duration of protection after one dose has not been fully established.18

Varicella vaccination is also strongly recommended for: 7

- nonimmune adolescents
- adults at high risk of exposure to chickenpox, such as healthcare workers and childcare workers

- nonimmune women before pregnancy
- parents
- nonimmune household contacts of immunocompromised people (to minimise the risk of VZV transmission to the susceptible person).

In the absence of contraindications, varicella vaccine can be given for postexposure prophylaxis within five days, preferably three days, of exposure to varicella to nonimmune age-eligible children and adults.14

Varicella vaccine contains live attenuated virus. It is contraindicated for people

TABLE. RECOMMENDED VARICELLA ZOSTER VIRUS VACCINE SCHEDULE ^{14,10,17}			
Recommended age	Vaccine	Recommended schedule	NIP funding
Children aged 12 months to less than 14 years	Measles, mumps, rubella, varicella (MMRV)	Single dose at 18 months	Yes
	Varicella	Single catch-up dose if routine schedule was missed	Yes
		Booster dose for improved protection	No
 Nonimmune adolescents 14 years and over Adults 	Varicella	Two doses, given at least four weeks apart	 Funded as per catch-up schedule to 19 years of age Free for all refugees and other humanitarian entrants¹⁶
Nonimmune household contact of people who are immunocompromised	Varicella	One or two doses depending on age	Depends on age (see above)
Adults 60 years and over	Zoster	Single dose	 Funded for people aged 70 years Free catch-up dose for people aged 71 to 79 years until October 2021¹⁷

PRACTICE POINTS ON VARICELLA FOR THE GP

- Varicella is a common and usually benign childhood illness that can nevertheless have serious complications in neonates, adults and people who are immunocompromised.
- Clinical manifestations develop 10 to 21 days after exposure to varicella zoster virus (or up to 28 days if the patient has received zoster immunoglobulin) and typically include a prodrome of fever and general malaise followed by development of a generalised vesicular rash.
- Varicella vaccine is a live attenuated vaccine that is generally safe and effective in preventing moderate to severe varicella.^{8,9}
- Varicella vaccination also reduces the risk of shingles in children.^{17,19}
- Two doses of varicella vaccine are recommended for primary vaccination, although only one dose is funded by the Australian National Immunisation Program and given in combination with measles, mumps and rubella vaccines (MMRV) at 18 months of age.
- Nonimmune adolescents aged 14 years or over and adults require two doses of varicella vaccine to achieve adequate protection.¹⁴
- Treatment of varicella in most children is supportive. Aspirin should be avoided because of the risk of Reye syndrome. Antiviral therapy is used for those at risk of severe disease.
- Any neonate, pregnant woman or immunocompromised person exposed to chickenpox should be urgently reviewed for consideration of aciclovir or zoster immunoglobulin prophylaxis.

who have had anaphylaxis to a dose or any component of the varicella-containing vaccine, pregnant women and people who are immunocompromised.^{7,14} A detailed list of medical conditions and therapies considered to cause severe immunosuppression can be found in the *Australian Immunisation Handbook*.¹⁴

Zoster vaccine

A single-dose live attenuated zoster vaccine is available in Australia and recommended for immunocompetent adults aged 60 years and over.¹⁷ It is currently funded under the National Shingles Vaccination Program for people aged 70 years, with catch up for those aged 71 to 79 years (until October 2021).¹⁷ Zoster vaccine contains 14 times the amount of live attenuated VZV compared with that in varicella vaccine.¹⁷ It is estimated to prevent 41% of cases of shingles and two-thirds of cases of postherpetic neuralgia in people aged 70 to 79 years.¹⁷

A prevaccination checklist, available on the Department of Health website,

can help immunisation providers screen for contraindications to zoster vaccine. The checklist needs to be completed before administration of the vaccine (www.health.gov.au/news/ clinical-update-pre-vaccination-checklist-for-zostavax-administration).²⁰

Complications occur in about 1% of [varicella] cases and are more common in neonates, adults and individuals who are immunocompromised

Zoster immunoglobulin

Postexposure prophylaxis with zoster immunoglobulin (known as ZIG or VZIG) provides some short-term passive immunity to VZV. It is recommended for people who have substantially higher risk of complications from chickenpox, including:

 people who are severely immunocompromised

- susceptible pregnant women
- neonates, particular those who are preterm or whose mothers develop primary chickenpox within seven days before or up to 28 days after delivery.^{14,15}

Zoster immunoglobulin is most effective if given within 96 hours of exposure, but some benefit may be conferred if given up to 10 days after exposure.^{12,14} Further information on specific indications for zoster immunoglobulin can be found in the *Australian Immunisation Handbook*.¹⁴

Contact precautions

People with chickenpox should be excluded from childcare, preschool, school and work until all vesicles have dried or for at least five days after the eruption first appears. They should also avoid contact with high-risk susceptible people, such as neonates, pregnant women and immunocompromised individuals.⁴ In healthcare settings, contact and airborne precautions should be implemented for patients with chickenpox.¹²

Conclusion

Despite a substantial reduction in morbidity and mortality from varicella following universal childhood varicella vaccination in Australia, this infection continues to cause serious (and sometimes fatal) complications in neonates, adults and people who are immunocompromised.21 GPs and other community health providers have an important role in identifying people who have chickenpox or have been exposed to chickenpox, especially high-risk individuals who require counselling, prophylaxis and potentially referral to hospital for further management. Practice points on varicella for GPs are summarised in the Box. MT

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A list of references is included in the online version of this article (www.medicinetoday.com.au).

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