Appendix. Alternative approaches to treating herpes zoster and herpetic neuralgia

Modality	Effector	Study method (Number of HZ patients)	Reported efficacy of evaluated effector against HZ and/or herpetic neuralgia	Country ^a (Source)
	Undaria pinnatifida	Effect of extract of <i>U. pinnatifida</i> (GFS) in 21 patients with herpetic infections (n=2)	GFS inhibited herpes virus reactivation and increased healing rate of herpetic outbreaks	Australia (Cooper 2002) ¹
	TCM herbs (unspecified decoction)	Randomised to TCM vs. western medicine: 1994–1995 (NR)	TCM better than western medicine (TC 93.3% vs. 85.8%, effective 75.5% vs. 52.2%)	China (Chen 2007) ²
	Ganoderma lucidum	Effect of G. lucidum on acute HZ pain and PHN (n=4)	Decreased severe HZ pain and intractable PHN	Japan (Hijikata 1998)³
		Effect of formula containing <i>G. lucidum</i> on acute HZ pain (n=5)	Rapid pain relief in a few days, almost complete by 10 days, no PHN through 1 year follow-up	Japan (Hijikata 2005) ⁴
	Keishikajutsubuto & Bushi-matsu	Effect of two herbs in PHN aggravated by cold environment (n=15)	Treatment is a promising way to treat intractable PHN	Japan (Nakanishi 2012)⁵
Botanic	Keisi-bushi-to & Hachimi-jio-gan	Effect of oriental herbal medicines on HZ pain (n=8)	Pain symptoms relieved in 2/3 treated with Keisi-bushi-to and 4/5 with Hachimi-jio-gan granules	Japan (Aki 2002) ⁶
	Yokukansan	Effect of Yokukansan in HZ pain and PHN (NR)	Yokukansan more effective than conventional analgesics	Japan (Nakamura 2009) ⁷
	Sairei-to	Pain relief in HZ (NR)	Analgesic efficacy 76.8%	Japan (Shukuwa 2002)8
	Clinacanthus nutans	Systematic review and meta-analysis of 4 RCTs (n=126)	Relative risk of 3.2 for 3-day full crusting (IQR 0.97–10.58), suggests beneficial effect in HZ	Thailand (Kongkaew 2011) ⁹
		Randomised to <i>C. nutans</i> 5% extract vs. placebo (n=51)	Faster healing and pain reduction vs. placebo (P<0.01), with no side effects.	Thailand (Sangkitporn 1995) ¹⁰
		Randomised to <i>C. nutans</i> 5% cream vs. placebo (n=75)	Faster lesion healing (27.5% vs. 8.5%)	Thailand (Charuwichitratana 1996) ¹¹
	Radix isatidis	R. isatidis prophylaxis against HZ in multiple myeloma treated with bortezomib (n=30)	Higher incidence of VZV reactivation without prophylaxis (25.0% vs. 0.0%, P=0.003)	China (Guo 2011) ¹²
	Hyperbaric oxygen (HBO₂)	Randomised to HBO ₂ + medication vs. medication (n=68)	HBO₂ enhanced efficacy (97.2% vs. 81.3%, P<0.05), faster healing, lower PHN rate and less depression	China (Peng 2012) ¹³
Molecular	Vitamin C	Effect of IV vitamin C in refractory HZ (n=1)	Treatment immediately reduced pain	Korea (Byun 2011) ¹⁴
	Bee venom	Effect of bee venom in refractory PHN (n=1)	Self-reported pain score fell from 8/10 to 2/10, with no adverse effects or rebound through 1 year follow-up	Korea (Lee 2014) ¹⁵
	Intramuscular stimulation (IMS)	Effect of IMS in treating PHN (n=4)	Self-reported pain decreased by more than half	Korea (Jung 2010) ¹⁶
		Meta-analysis of 10 RCTs (NR)	Superior clinical cure and analgesia to western medicine (P<0.01), more high-quality studies needed	China (Yu 2007) ¹⁷
Mechanic	Acupuncture (AP)	Randomised to AP vs. ibuprofen + famotidine + IV vitamins $B_1 \& B_{12}$ (n=24)	Treatment superior (TE 91.6% vs. 83.3%)	China (Fang 2011) ¹⁸
		Case series of nerve cell puncturing (n=48)	Good therapeutic results	China (Ni 2008) ¹⁹
	Cupping	Meta-analysis of 135 RCTs:b 1992–2010 (NR)	Potential effect, but need more well designed trials	Australia (Cao 2012) ²⁰
		Systematic review of 73 RCTs, 22 trials, 373 case series, 82 cases: 1959–2008 (NR)	Potential benefit but more rigorously designed trials warranted	China (Cao 2010) ²¹

	Blood letting	Systematic review of 98,526 cases involving 261 diseases (NR)	Bleeding therapy has evident advantages for treating HZ	China (Chen 2009) ²²
		Case study of bleeding point ear apex (n=1)	Immediate relief of most pain, two treatments cured	China (Wang 2006) ²³
	Far-infrared radiation	Effect of far-infrared radiation in HZ (n=50)	Superior healing and pain relief vs. control	China (Zhang 1994) ²⁴
Energetic	CT-guided radiofrequency (RF) thermocoagulation	Effect of CT-guided RF dorsal root ganglion thermocoagulation in intractable PHN (n=16)	Hyperalgesia disappeared, immediate self-reported pain scores fell from 7–9/10 to 2–3/10, with no relapse	China (Huang 2008) ²⁵
	Laser	Effect of combi laser therapy in PHN (n=50)	Cure 88.0% after 20 exposures, relief 10% included four HZ ophthalmicus cases	India (Mittal 1996) ²⁶
		Case-control study of adding He-Ne laser to medication in HZ keratitis (NR)	Laser enhanced efficacy with accelerated healing and better pain relief (P<0.05)	China (Li 2012) ²⁷
	Temporary electric spinal cord stimulation (SCS)	Effect of adding SCS to epidural block in refractory PHN (n=14)	SCS reduced self-reported pain and spared epidural anaesthesia	Japan (Moriyama 2009) ²⁸
	Light-emitting diode (LED)	Randomised to LED + IV acyclovir + analgesia vs. IV acyclovir + analgesia (n=60)	LED accelerated relief of acute pain (1.4 vs. 1.8 weeks, P<0.05) but had no effect on onset of PHN	Korea (Jung 2013) ²⁹
		HZ ophthalmicus patients randomised to 830 nm LED vs. oral famciclovir (n=28)	LED accelerated healing (13.1 days vs. 15.9 days, P=0.006) and reduced self-reported pain from day 4	Korea (Park 2013) ³⁰
	Ultrasound (US)-guided pulsed RF	Effect of US-guided pulsed RF in refractory facial HZ (n=1)	Pain reduced, persisting through 6-months follow-up, with no need for further management	Korea (Lim 2013) ³¹
	Scrambler	Effect of scrambler therapy in refractory PHN (n=3)	Self-reported pain decreased by 50% after 10 sessions	Korea (Ko 2013) ³²
	AP + blood-letting with cupping & surround needling	Randomised to combination vs. oral + topical aciclovir (n=240)	Combination more effective (92.5% vs. 55.8%, P<0.01), with more rapid pain relief and healing	China (Huo. 2007) ³³
	Acupoint selection + AP + cupping	Randomised to combination at differential acupoints vs. affected areas (n=86)	Targeted-therapy superior (cure & effective 93.0% vs. 67.4%, P<0.01), lower PHN rate (2.3% vs. 14.0%)	China (Pan 2011) ³⁴
	AP + electro-AP, moxibustion hot needle, tapping + cupping	Randomised to various mechanic TCM modalities vs. western medicine (n=500)	No significant difference in efficacy, but TCM more cost- effective	China (Li 2012) ³⁵
	Electro AP + AP or cotton moxibustion or hot needle or tapping/cupping	Randomised to various TCM modalities vs. oral valaciclovir + vitamin B_1 (n=189)	No significant differences in clinical cure efficacy but TCM significantly reduced pain from post-treatment day 7 (P<0.05)	China (Huang 2012) ³⁶
Combined	Collateral-pricking + 'healthy energy strengthening'	Retrospective comparison of collateral pricking vs. aciclovir (n=66)	Treatment superior (cure 95.6% vs. 75.6%, P<0.05)	China (Yu 2005) ³⁷
	Needling + moxibustion	Randomised to moxibustion at area of surround needling vs. needling only (n=72)	Treatment more effective (97.4% vs. 85.3%, P<0.05), faster healing and reduced residual neuralgia	China (Zhang 2007) ³⁸
	AP + point injection	Effect of combination in PHN (n=30)	Combination has curative effect on PHN (TE 90%)	China (Wang 2008) ³⁹
	Needling + thin-cotton moxibustion	Randomised to combination vs. oral + topical aciclovir (n=80)	Treatment superior (effective 78.6% vs. 39.5%, P<0.05)	China (Tian 2011) ⁴⁰
	Plum-blossom needle tapping + moxibustion	Randomised to combination vs. topical aciclovir + oral valaciclovir + vitamin B_{12} (n=120)	Treatment superior (cure 80.0% vs. 45.0%, P<0.10; TE 98.3% vs. 71.7%, P<0.05) and faster, recurrence 1.6% vs. 13.3%	China (Yang 2012) ⁴¹
	Percussopunctato + cupping	Effect of combination in HZ and PHN (n=33)	TE 100%: cure 60.6%, good effect 33.3%, effect 6%	China (Han 2003) ⁴²

Ear tapping & pressing + acupoint vitamin B ₁₂ injection	Randomised to combination vs. standard western medicine (n=116)	Cure 60% vs. 28.6%, effective 83.3% vs. 50.0%, (P<0.05), faster pain relief and cessation (P<0.01)	China (Wu 2007) ⁴³
Blood-letting & cupping + UV irradiation	Randomised to combination vs. standard therapy including aciclovir (n=130)	Higher and faster cure & effective rates (respectively, 76.9% vs. 38.5% & 90.8% vs. 66.2%, P<0.01), lower PHN rate (3.1% vs. 12.3%, P<0.05)	China (Ouyang 2009)
Electro-AP + needling	Randomised to combination vs. valaciclovir + vitamin B_1 (n=80)	Combination superior (cure 75.0% vs. 37.5%, effective 92.5% vs. 67.5, with faster healing (P<0.01)	China (Li 2009) ⁴⁵
Electro-AP + blood-letting & cupping	Randomised to combination vs. valaciclovir + indomethacin + vitamins $B_1 \& B_{12}$ (n=53)	Cure & effective 96.8% vs. 81.8% (P<0.05)	China (Liu 2009) ⁴⁶
Electro-AP + collateral pricking & cupping	Randomised to combination vs. oral valaciclovir + indomethacin (n=53)	Combination superior (cure & effective 93.5% vs. 72.7%, P<0.01)	China (Liu 2013) ⁴⁷
AP & cupping + blood letting	Case report (n=1)	One treatment reduced pain and five cured HZ, with no residual pain or sequelae at 3 months	China (Hu 2001) ⁴⁸
Electronic meridian detection + blood-letting & cupping	Randomised to combination vs. oral acyclovir (n=200)	Treatment superior (TE 100% vs. 60%, P<0.001), faster pain relief and healing, PHN rate 0.0% vs. 26%	China (Dai 2011) ⁴⁹
Variable frequency electro-AP + herbal moxibustion	Randomised to combination vs. ibuprofen + vitamins B_1 & E (n=37)	Treatment superior to medication (effective 94.44% vs. 73.68%, P<0.05)	China (Wang 2012) ⁵⁰
Swift needling + medication	Randomised to fire-AP + oral valaciclovir + vitamin B_1 vs. oral valaciclovir + vitamin B_1 (n=79)	Combination superior (TE 95.1% vs. 89.5%, P<0.05)	China (Fan 2013) ⁵¹
Subcutaneous needling + laser	PHN patients randomised to electro-AP + laser vs. electro-AP (n=102)	Combination superior (TE 94.1% vs. 68.6%, recovery 56.9% vs. 35.3%, P<0.05), with faster action and cure and less relapse (6.9% vs. 44.4%, P<0.05)	China (Guo 2014) ⁵²
Encircled AP + valaciclovir	Effect of encircled AP + oral valaciclovir vs. oral valaciclovir (n=60)	Combination superior, faster healing and pain relief	China (He 2007) ⁵³
Surround AP + He-Ne laser	Randomised to surround AP + IV aciclovir + He-Ne laser vs. IV & topical aciclovir (n=60)	Combination superior in effective rate, healing, pain relief and time to cure (P<0.05)	China (Song 2013) ⁵⁵⁴
TCM + mecobalamin	PHN patients randomised to mecobalamin or TCM vs, mecobalamin + TCM (n=160)	Combination superior (effective 90.9% vs. 72% mecobalamin & 70.9% TCM, P<0.01)	China (Tian 2005) ⁵⁵
Linear polarised near-infrared light (LPIR) + fentanyl patch	Randomised to combination vs. LPIR or fentanyl patch (n=30)	Combination superior (80% vs. 60% & 70%, P<0.05), self-reported pain fell from baseline (P<0.05)	China (Wang 2007) ⁵⁶
Electro-AP + herbal medicine	Randomised to combination vs. oral + topical aciclovir + mecobalamin (n=62)	Combination superior (P<0.01)	China (Wang 2012) ⁵⁷
Oral TCM + AP, moxibustion, corticosteroids, antivirals, analgesia and trophic nerve	Randomised to combination vs. conventional antiviral and analgesic agents ± corticosteroids ± AP & moxibustion (n=128)	Composite treatment with oral TCM significantly eased self-reported pain and improved quality of life	China (Xie 2003) ⁵⁸

AP acupuncture, HZ herpes zoster, NR not reported, UV ultraviolet, PHN post-herpetic neuralgia, HBO₂ hyperbaric oxygen, RCT randomised controlled trial, TCM traditional Chinese medicine, LED light-emitting diode, IV intravenous, US ultrasound, IQR interquartile range, TC total cure, TE total effective, RF radiofrequency, CT computed tomography, SCS spinal cord stimulation.

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^b Included cupping types: wet, retained, moving, flash, medicinal, needle, combined.

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