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Non - neurotoxic activity of Malayan krait (*Bungarus candidus*) venom from Thailand (Article)

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Abstract

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Background: Envenoming by kraits (genus *Bungarus*) is a medically significant issue in South Asia and Southeast Asia.

Malayan krait (*Bungarus candidus*) venom is known to contain highly potent neurotoxins. In recent years, there have been reports on the non - neurotoxic activities of krait venom that include myotoxicity and nephrotoxicity. However, research on such non -neurotoxicity activities of Malayan krait venom is extremely limited. Thus, the aim of the present study was to determine the myotoxic, cytotoxic and nephrotoxic activities of *B. candidus* venoms from northeastern (BC-NE) and southern (BC-S) Thailand in experimentally envenomed rats. Methods: Rats were administered Malayan krait (BC-NE or BC-S) venom (50 µg/kg, i.m.) or 0.9% NaCl solution (50 µL, i.m.) into the right hind limb. The animals were sacrificed 3, 6 and 24 h after venom administration. The right gastrocnemius muscle and both kidneys were collected for histopathological analysis. Blood samples were also taken for determination of creatine kinase (CK) and lactate dehydrogenase (LDH) levels. The human embryonic kidney cell line (HEK-293) was used in a cell proliferation assay to determine cytotoxic activity. Results: Administration of BC-NE or BC-S venom (50 µg/kg, i.m.) caused time-dependent myotoxicity, characterized by an elevation of CK and LDH levels. Histopathological examination of skeletal muscle displayed marked muscle necrosis and myofiber disintegration 24 h following venom administration. Both Malayan krait venoms also induced extensive renal tubular injury with glomerular and interstitial congestion in rats. BC-NE and BC-S venoms (100-0.2 µg/mL) caused concentration-dependent cytotoxicity on the HEK-293 cell line. However, BC-NE venom (IC₅₀ = 8 ± 1 µg/mL; at 24 h incubation; n = 4) was found to be significantly more cytotoxic than BC-S venom (IC₅₀ = 15 ± 2 µg/mL; at 24 h incubation; n = 4). In addition, the PLA₂ activity of BC-NE venom was significantly higher than that of BC-S venom. Conclusions: This study found that Malayan krait venoms from both populations possess myotoxic, cytotoxic and nephrotoxic activities. These findings may aid in clinical diagnosis and treatment of envenomed patients in the future. © 2018 The Author(s).

Author keywords

[Bungarus candidus](#) [Kidney](#) [Krait](#) [Myotoxicity](#) [Nephrotoxicity](#) [Venom](#)

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Chaisakul, J. , Parkington, H.C. , Isbister, G.K.

(2013) *Basic and Clinical Pharmacology and Toxicology*Pharmacokinetics and pharmacodynamics of the myotoxic venom of *Pseudechis australis* (mulga snake) in the anesthetised rat

Hart, A.J. , Hodgson, W.C. , O'leary, M.

(2014) *Clinical Toxicology*

EMTREE drug terms:

chloride ion creatine kinase creatinine lactate dehydrogenase nitrogen
phospholipase A2 potassium ion snake venom sodium ion urea

EMTREE medical terms:

animal experiment animal tissue Article blood sampling Bungarus
Bungarus candidus cell proliferation assay cell viability assay chloride blood level
concentration (parameters) controlled study creatine kinase blood level cytotoxicity
embryo envenomation enzyme activity enzyme assay gastrocnemius muscle
geographic distribution HEK293 cell line histopathology human human cell IC50
kidney injury kidney tubule disorder lactate dehydrogenase blood level male
muscle disease muscle necrosis myotoxicity nephrotoxicity nonhuman
potassium blood level rat skeletal muscle sodium blood level Thailand toxicity
urea nitrogen blood level

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- 1 *Guidelines for the management of snake-bites*. Cited 2 times.
In: 2nd edition; 2016.

- 2 Rusmili, M.R.A., Yee, T.T., Mustafa, M.R., Hodgson, W.C., Othman, I.
Proteomic characterization and comparison of Malaysian Bungarus candidus and Bungarus fasciatus venoms

(2014) *Journal of Proteomics*, 110, pp. 129-144. Cited 19 times.

<http://www.elsevier.com>

doi: 10.1016/j.jprot.2014.08.001

[View at Publisher](#)

- 3 Nirthanan, S., Charpantier, E., Gopalakrishnakone, P., Gwee, M.C.E., Khoo, H.E., Cheah, L.S., Kini, R.M., (...), Bertrand, D.
Neuromuscular effects of candoxin, a novel toxin from the venom of the Malayan krait (Bungarus candidus)

(2003) *British Journal of Pharmacology*, 139 (4), pp. 832-844. Cited 30 times.

doi: 10.1038/sj.bjp.0705299

[View at Publisher](#)

- 4 Trinh, K.X., Khac, Q.L., Trinh, L.X., Warrell, D.A.
Hyponatraemia, rhabdomyolysis, alterations in blood pressure and persistent mydriasis in patients envenomed by Malayan kraits (Bungarus candidus) in southern Viet Nam

(2010) *Toxicon*, 56 (6), pp. 1070-1075. Cited 19 times.

doi: 10.1016/j.toxicon.2010.06.026

[View at Publisher](#)

- 5 Chaisakul, J., Ahmad Rusmili, M.R., Hodgson, W.C., Hatthachote, P., Suwan, K., Inchan, A., Chanhome, L., (...), Chootip, K.

A pharmacological examination of the cardiovascular effects of Malayan krait (*Bungarus candidus*) venoms

(2017) *Toxins*, 9 (4), art. no. 122. Cited 2 times.
<http://www.mdpi.com/2072-6651/9/4/122/pdf>
doi: 10.3390/toxins9040122

[View at Publisher](#)

- 6 Hart, A.J., Hodgson, W.C., O'leary, M., Isbister, G.K.

Pharmacokinetics and pharmacodynamics of the myotoxic venom of *Pseudechis australis* (mulga snake) in the anesthetised rat

(2014) *Clinical Toxicology*, 52 (6), pp. 604-610. Cited 3 times.
<http://www.informahealthcare.com>
doi: 10.3109/15563650.2014.914526

[View at Publisher](#)

- 7 Azevedo-Marques, M.M., Hering, S.E., Cupo, P.

Evidence that *Crotalus durissus terrificus* (South American rattlesnake) envenomation in humans causes myolysis rather than hemolysis

(1987) *Toxicon*, 25 (11), pp. 1163-1168. Cited 62 times.
doi: 10.1016/0041-0101(87)90134-6

[View at Publisher](#)

- 8 Ali, S.A., Alam, J.M., Abbasi, A., Zaidi, Z.H., Stoeva, S., Voelter, W.

Sea snake *Hydrophis cyanocinctus* venom. II. Histopathological changes, induced by a myotoxic phospholipase A₂ (PLA₂-H1)

(2000) *Toxicon*, 38 (5), pp. 687-705. Cited 19 times.
doi: 10.1016/S0041-0101(99)00184-1

[View at Publisher](#)

- 9 Gutiérrez, J.M., Alberto Ponce-Soto, L., Marangoni, S., Lomonte, B.

Systemic and local myotoxicity induced by snake venom group II phospholipases A₂: Comparison between crotoxin, crotoxin B and a Lys49 PLA₂ homologue

(2008) *Toxicon*, 51 (1), pp. 80-92. Cited 51 times.
doi: 10.1016/j.toxicon.2007.08.007

[View at Publisher](#)

- 10 Melo, P.A., Burns, C.F., Blankemeyer, J.T., Ownby, C.L.

Membrane depolarization is the initial action of crotoxin on isolated murine skeletal muscle

(2004) *Toxicon*, 43 (2), pp. 111-119. Cited 14 times.
doi: 10.1016/j.toxicon.2003.10.029

[View at Publisher](#)

- 11 Mora-Obando, D., Díaz, C., Angulo, Y., Gutiérrez, J.M., Lomonte, B.

Role of enzymatic activity in muscle damage and cytotoxicity induced by *Bothrops asper* Asp49 phospholipase A₂ myotoxins: Are there additional effector mechanisms involved?

(2014) *PeerJ*, 2014 (1), art. no. e569. Cited 10 times.
<https://peerj.com/articles/569.pdf>
doi: 10.7717/peerj.569

[View at Publisher](#)

- 12 Hart, A.J., Smith, A.I., Reeve, S., Hodgson, W.C.
Isolation and characterisation of acanmyotoxin-2 and acanmyotoxin-3, myotoxins from the venom of the death adder *Acanthophis* sp. *Seram*
(2005) *Biochemical Pharmacology*, 70 (12), pp. 1807-1813. Cited 13 times.
doi: 10.1016/j.bcp.2005.09.017
[View at Publisher](#)
-
- 13 Chaisakul, J., Parkington, H.C., Isbister, G.K., Konstantakopoulos, N., Hodgson, W.C.
Differential myotoxic and cytotoxic activities of pre-synaptic neurotoxins from papuan taipan (*Oxyuranus scutellatus*) and irian jayan death adder (*Acanthophis rugosus*) venoms
(2013) *Basic and Clinical Pharmacology and Toxicology*, 112 (5), pp. 325-334. Cited 3 times.
doi: 10.1111/bcpt.12048
[View at Publisher](#)
-
- 14 Warrell, D.A., Looareesuwan, S., White, N.J., Theakston, R.D., Warrell, M.J., Kosakarn, W., Reid, H.A.
Severe neurotoxic envenoming by the Malayan krait *Bungarus candidus* (Linnaeus): Response to antivenom and anticholinesterase
(1983) *British Medical Journal*, 286 (6366), pp. 678-680. Cited 79 times.
[View at Publisher](#)
-
- 15 Skejić, J., Hodgson, W.C.
Population Divergence in Venom Bioactivities of Elapid Snake *Pseudonaja textilis*: Role of Procoagulant Proteins in Rapid Rodent Prey Incapacitation ([Open Access](#))
(2013) *PLoS ONE*, 8 (5), art. no. e63988. Cited 14 times.
<http://www.plosone.org/article/fetchObjectAttachment.action?uri=info%3Adoi%2F10.1371%2Fjournal.pone.0063988&representation=PDF>
doi: 10.1371/journal.pone.0063988
[View at Publisher](#)
-
- 16 Wickramaratna, J.C., Fry, B.G., Hodgson, W.C.
Species-dependent variations in the vitro myotoxicity of death adder (*Acanthophis*) venoms
(2003) *Toxicological Sciences*, 74 (2), pp. 352-360. Cited 24 times.
doi: 10.1093/toxsci/kfg144
[View at Publisher](#)
-
- 17 Vongsak, B., Mangmool, S., Gritsanapan, W.
Antioxidant Activity and Induction of mRNA Expressions of Antioxidant Enzymes in HEK-293 Cells of *Moringa oleifera* Leaf Extract
(2015) *Planta Medica*, 81 (12-13), pp. 1084-1089. Cited 13 times.
<http://www.thieme-connect.com/ejournals/toc/plantamedica>
doi: 10.1055/s-0035-1546168
[View at Publisher](#)
-
- 18 Chaisakul, J., Konstantakopoulos, N., Smith, A.I., Hodgson, W.C.
Isolation and characterisation of P-EPTX-Ap1a and P-EPTX-Ar1a: Pre-synaptic neurotoxins from the venom of the northern (*Acanthophis praelongus*) and Irian Jayan (*Acanthophis rugosus*) death adders
(2010) *Biochemical Pharmacology*, 80 (6), pp. 895-902. Cited 15 times.
doi: 10.1016/j.bcp.2010.05.008
[View at Publisher](#)

- 19 Silva, A., Johnston, C., Kuruppu, S., Kneisz, D., Maduwage, K., Kleifeld, O., Smith, A.I., (...), Isbister, G.K.
Clinical and Pharmacological Investigation of Myotoxicity in Sri Lankan Russell's Viper (*Daboia russelii*) Envenoming ([Open Access](#))
- (2016) *PLoS Neglected Tropical Diseases*, 10 (12), art. no. e0005172.
<http://www.plosntds.org/index.php>
doi: 10.1371/journal.pntd.0005172
- [View at Publisher](#)
-
- 20 Johnston, C.I., Brown, S.G.A., O'Leary, M.A., Currie, B.J., Greenberg, R., Taylor, M., Barnes, C., (...), Isbister, G.K.
Mulga snake (*Pseudechis australis*) envenoming: A spectrum of myotoxicity, anticoagulant coagulopathy, haemolysis and the role of early antivenom therapy- Australian Snakebite Project (ASP-19)
- (2013) *Clinical Toxicology*, 51 (5), pp. 417-424. Cited 19 times.
doi: 10.3109/15563650.2013.787535
- [View at Publisher](#)
-
- 21 Denis, D., Lamireau, T., Llanas, B., Bedry, R., Fayon, M.
Rhabdomyolysis in European viper bite
- (1998) *Acta Paediatrica, International Journal of Paediatrics*, 87 (9), pp. 1013-1015. Cited 11 times.
doi: 10.1080/080352598750031743
- [View at Publisher](#)
-
- 22 Pandey, D.P., Sharma, S.K., Alirol, E., Chappuis, F., Kuch, U.
Fatal neurotoxic envenomation following the bite of a greater black krait (*Bungarus niger*) in Nepal: A case report
- (2016) *Journal of Venomous Animals and Toxins Including Tropical Diseases*, 22 (1), art. no. 19.
<http://www.jvat.org/>
doi: 10.1186/s40409-016-0073-8
- [View at Publisher](#)
-
- 23 Faiz, M.A., Ghose, A., Ahsan, M.F., Rahman, M.R., Amin, M.R., Hassan, M.M.U., Chowdhury, M.A.W., (...), Warrell, D.A.
The greater black krait (*Bungarus niger*), a newly recognized cause of neuro-myotoxic snake bite envenoming in Bangladesh
- (2010) *Brain*, 133 (11), pp. 3181-3193. Cited 29 times.
<http://brain.oxfordjournals.org/>
doi: 10.1093/brain/awq265
- [View at Publisher](#)
-
- 24 Pochanugool, C., Wilde, H., Jitapunkul, S., Limthongkul, S.
Spontaneous recovery from severe neurotoxic envenoming by a Malayan krait *Bungarus candidns* (linnaeus) in Thailand
- (1997) *Wilderness and Environmental Medicine*, 8 (4), pp. 223-225. Cited 15 times.
- [View at Publisher](#)
-
- 25 De Roodt, A.R., Lago, N.R., Stock, R.P.
Myotoxicity and nephrotoxicity by *Micrurus* venoms in experimental envenomation
- (2012) *Toxicon*, 59 (2), pp. 356-364. Cited 13 times.
doi: 10.1016/j.toxicon.2011.11.009
- [View at Publisher](#)

- 26 Ponraj, D., Gopalakrishnakone, P.
Morphological changes induced by a generalized myotoxin (myoglobinuria-inducing toxin) from the venom of *Pseudechis australis* (king brown snake) in skeletal muscle and kidney of mice
(1995) *Toxicon*, 33 (11), pp. 1453-1467. Cited 16 times.
doi: 10.1016/0041-0101(95)00091-Y
[View at Publisher](#)
-
- 27 Sitprija, V.
Snakebite nephropathy (review article)
(2006) *Nephrology*, 11 (5), pp. 442-448. Cited 56 times.
doi: 10.1111/j.1440-1797.2006.00599.x
[View at Publisher](#)
-
- 28 Mebs, D., Ownby, C.L.
Myotoxic components of snake venoms: Their biochemical and biological activities
(1990) *Pharmacology and Therapeutics*, 48 (2), pp. 223-236. Cited 158 times.
doi: 10.1016/0163-7258(90)90081-C
[View at Publisher](#)
-
- 29 Gasanov, S.E., Dagda, R.K., Rael, E.D.
Snake venom cytotoxins, phospholipase A₂s, and Zn²⁺-dependent metalloproteinases: mechanisms of action and pharmacological relevance
(2014) *J Clin Toxicol*, 4 (1). Cited 27 times.
-
- 30 Chaiyabutr, N., Vasaruchapong, T., Chanhome, L., Rungsipipat, A., Sitprija, V.
Acute effect of Russell's viper (*Daboia siamensis*) venom on renal tubular handling of sodium in isolated rabbit kidney ([Open Access](#))
(2014) *Asian Biomedicine*, 8 (2), pp. 195-202. Cited 2 times.
<http://abm.digitaljournals.org/index.php/abm/article/viewFile/2482/833>
doi: 10.5372/1905-7415.0802.279
[View at Publisher](#)
-
- 31 Ponraj, D., Gopalakrishnakone, P.
Renal lesions in rhabdomyolysis caused by *Pseudechis australis* snake myotoxin ([Open Access](#))
(1997) *Kidney International*, 51 (6), pp. 1956-1969. Cited 24 times.
<https://www.journals.elsevier.com/kidney-international>
doi: 10.1038/ki.1997.267
[View at Publisher](#)
-
- 32 Ponraj, D., Gopalakrishnakone, P.
Establishment of an animal model for myoglobinuria by use of a myotoxin from *Pseudechis australis* (King Brown Snake) venom in mice
(1996) *Laboratory Animal Science*, 46 (4), pp. 393-398. Cited 26 times.
-
- 33 Lomonte, B., Rangel, J.
Snake venom Lys49 myotoxins: From phospholipases A₂ to non-enzymatic membrane disruptors
(2012) *Toxicon*, 60 (4), pp. 520-530. Cited 52 times.
doi: 10.1016/j.toxicon.2012.02.007
[View at Publisher](#)

- 34 Oguiura, N., Boni-Mitake, M., Rádis-Baptista, G.
New view on crotamine, a small basic polypeptide myotoxin from South American rattlesnake venom

(2005) *Toxicon*, 46 (4), pp. 363-370. Cited 43 times.
doi: 10.1016/j.toxicon.2005.06.009

[View at Publisher](#)

- 35 Rádis-Baptista, G., Kerkis, I.
Crotamine, a small basic polypeptide myotoxin from rattlesnake venom with cell-penetrating properties.

(2011) *Current pharmaceutical design*, 17 (38), pp. 4351-4361. Cited 23 times.

[View at Publisher](#)

- 36 Guardiola, O., Andolfi, G., Tirone, M., Iavarone, F., Brunelli, S., Minchiotti, G.
Induction of acute skeletal muscle regeneration by cardiotoxin injection

(2017) *Journal of Visualized Experiments*, 2017 (119), art. no. e54515. Cited 2 times.
<http://www.jove.com/pdf/54515/jove-protocol-54515-induction-acute-skeletal-muscle-regeneration-cardiotoxin>
doi: 10.3791/54515

[View at Publisher](#)

- 37 Kini, R.M., Doley, R.
Structure, function and evolution of three-finger toxins: Mini proteins with multiple targets

(2010) *Toxicon*, 56 (6), pp. 855-867. Cited 143 times.
doi: 10.1016/j.toxicon.2010.07.010

[View at Publisher](#)

- 38 Salvador, G.H.M., dos Santos, J.I., Lomonte, B., Fontes, M.R.M.
Crystal structure of a phospholipase A₂ from *Bothrops asper* venom: Insights into a new putative "myotoxic cluster"

(2017) *Biochimie*, 133, pp. 95-102.
www.elsevier.com/locate/biochimie
doi: 10.1016/j.biochi.2016.12.015

[View at Publisher](#)

- 39 Kalam, Y., Isbister, G.K., Mirtschin, P., Hodgson, W.C., Konstantakopoulos, N.
Validation of a cell-based assay to differentiate between the cytotoxic effects of elapid snake venoms

(2011) *Journal of Pharmacological and Toxicological Methods*, 63 (2), pp. 137-142. Cited 13 times.
doi: 10.1016/j.vascn.2010.09.001

[View at Publisher](#)

- 40 Jamunaa, A., Vejayan, J., Halijah, I., Sharifah, S.H., Ambu, S.
Cytotoxicity of Southeast Asian snake venoms

(2012) *Journal of Venomous Animals and Toxins Including Tropical Diseases*, 18 (2), pp. 150-156. Cited 8 times.
<http://www.scielo.br/pdf/jvatiid/v18n2/v18n2a04.pdf>
doi: 10.1590/S1678-91992012000200004

[View at Publisher](#)

□ 41 Winter, K.L., Isbister, G.K., McGowan, S., Konstantakopoulos, N., Seymour, J.E., Hodgson, W.C.

A pharmacological and biochemical examination of the geographical variation of *Chironex fleckeri* venom

(2010) *Toxicology Letters*, 192 (3), pp. 419-424. Cited 18 times.
doi: 10.1016/j.toxlet.2009.11.019

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