

Material and Methods

We conducted a systematic search, for this meta-summary, from the databases of PubMed, and Science Direct. The search terms included ("drug overdose" OR poisoning OR hypoglycemia OR "lactic acidosis" AND Metformin).

Further, it was filtered for the case reports published in the English language, and on adult (> 18 years) humans. We screened all the search results manually, and we included the relevant literature for Metformin toxicity only. Duplicate articles from different search databases were excluded manually.

Data analysis

We prepared the datasheet and evaluated it with help of Excel, Microsoft office 2019. Categorical variables were presented as frequency and percentage. Mean [standard deviation (SD)] or median [interquartile range (IQR)] was used for continuous variables as appropriate. We applied a non-parametric correlational statistical test to test the non-parametric statistical hypothesis as found appropriate. A p-value of < 0.05 was deemed significant.

All the statistical analyses were done using SPSS (version 25.0, IBM SPSS Inc., Chicago, IL, United States) unless otherwise indicated. Tabulation and final documentation were done using MS Office software (MS office 2019, Microsoft Corp, WA, United States).

Supplementary Table 1 Literature search

Database	PubMed	Science direct	Google scholar
keywords	("drug overdose" OR "adverse drug reaction" OR poisoning OR hypoglycemia OR "lactic acidosis" OR "metabolic acidosis") AND Metformin	("drug overdose" OR "adverse drug reaction" OR poisoning OR hypoglycemia OR "lactic acidosis" OR "metabolic acidosis") AND Metformin	("drug overdose" OR "adverse drug reaction" OR poisoning OR hypoglycemia OR hypoglycemia OR "lactic acidosis" OR "lactic acidosis" OR "metabolic acidosis") AND Metformin AND "Case Report"
Page link	https://pubmed.ncbi.nlm.nih.gov/?term=%28%22drug+overdose%22+OR+%22adverse+drug+reaction%22+OR+poisoning+OR+hypoglycemia+OR+%22lactic+acidosis%22+OR+%22metabolic+acidosis%22%29+AND+Metformin&filt	https://www.sciencedirect.com/search?q=%28%22drug%20overdose%22%20OR%20%22adverse%20drug%20reaction%22%20OR%20poisoning%20OR%20hypoglycemia%20OR%20%22lactic%20acidosis%22%20OR%20%22metabolic%20acidosis%22%29%2	https://scholar.google.com/scholar?hl=en&as_sdt=0%2C5&q=%28%22drug+overdose%22+OR+%22adverse+drug+reaction%22+OR+poisoning+OR+hypoglycemia+OR+lactic+acidosis%22+OR+%22metabolic+acidosis%22%29+AND+Metformin+AND+Case+Report

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Primary results (n) (keyword results)	3590	10856	76800
Screening (n) (after applying filters)	291	149	12800
Eligibility (by sorting manually the relevance of each article)(n)	263	27	
Inclusion (n) (manually sorted list of final literature set after removal of duplicates)	184		

Supplementary Table 2 Counts of studies and cases

	Counts of studies	Counts of cases
Total case reports	158	158
Total case series	26	84

1. Elshafei MN, Alamin M, Mohamed MFH. Osmolar-gap in the setting of metformin-associated lactic acidosis: Case report and a literature review highlighting an apparently unusual association. *Medicine* [Internet]. 2020 Oct 9 [cited 2022 Jan 18];99(41):e22492. Available from: <https://journals.lww.com/10.1097/MD.00000000000022492>
2. McCabe DJ, Baker S, Stellpflug SJ. Hemodialysis in metformin-associated lactic acidosis due to acute overdose in a metformin-naïve patient. *The American Journal of Emergency Medicine* [Internet]. 2018 Sep [cited 2022 Jan 18];36(9):1721.e1-1721.e2. Available from: <https://linkinghub.elsevier.com/retrieve/pii/S0735675718304522>
3. Fadden EJ, Longley C, Mahambrey T. Metformin-associated lactic acidosis. *BMJ Case Rep* [Internet]. 2021 Jul [cited 2022 Jan 18];14(7):e239154. Available from: <https://casereports.bmj.com/lookup/doi/10.1136/bcr-2020-239154>
4. Sahutoglu T, Sahutoglu E. Hemodialysis during cardiopulmonary resuscitation in patients with severe metabolic acidosis. *Int J Artif Organs* [Internet]. 2018 Aug [cited 2022 Jan 18];41(8):431–6. Available from: <http://journals.sagepub.com/doi/10.1177/0391398818784275>
5. Suzuki K, Okada H, Yoshida S, Okamoto H, Suzuki A, Suzuki K, et al. Effect of high-flow high-volume-intermittent hemodiafiltration on metformin-associated lactic acidosis with circulatory failure: a case report. *J Med Case Reports* [Internet]. 2018 Dec [cited 2022 Jan 18];12(1):280. Available from:

<https://jmedicalcasereports.biomedcentral.com/articles/10.1186/s13256-018-1809-6>

6. Tirelli F, Biondi A, Persiani R. Fatal Case of Metformin-Associated Lactic Acidosis Associated With Temporary Ileostomy: A Case Report. *Journal of Wound, Ostomy & Continence Nursing* [Internet]. 2018 Jul [cited 2022 Jan 18];45(4):364–5. Available from: <https://journals.lww.com/00152192-201807000-00014>
7. Chen T, Zhu C, Liu B. Extracorporeal membrane oxygenation with continuous renal replacement therapy to treat metformin-associated lactic acidosis: A case report. *Medicine* [Internet]. 2020 Jun 26 [cited 2022 Jan 18];99(26):e20990. Available from: <https://journals.lww.com/10.1097/MD.00000000000020990>
8. Goonoo MS, Morris R, Raithatha A, Creagh F. Metformin-associated lactic acidosis: reinforcing learning points. *BMJ Case Rep* [Internet]. 2020 Sep [cited 2022 Jan 18];13(9):e235608. Available from: <https://casereports.bmj.com/lookup/doi/10.1136/bcr-2020-235608>
9. Aldobeaban S, Mzahim B, Alshehri AA. Recurrent hypoglycemia secondary to metformin toxicity in the absence of co-ingestions: a case report. *J Med Case Reports* [Internet]. 2018 Dec [cited 2022 Jan 18];12(1):223. Available from: <https://jmedicalcasereports.biomedcentral.com/articles/10.1186/s13256-018-1758-0>
10. Ali S, Labuschagne H, Azarov N, Hindi Z, Oud L. Metformin-associated lactic acidosis mimicking ischaemic bowel. *BMJ Case Reports* [Internet]. 2018 Feb 1 [cited 2022 Jan 18];bcr-2017-221686. Available from: <https://casereports.bmj.com/lookup/doi/10.1136/bcr-2017-221686>
11. Chiew AL, Wright DFB, Dobos NM, McArdle K, Mostafa AA, Newth A, et al. ‘Massive’ metformin overdose: ‘Massive’ metformin overdose. *Br J Clin Pharmacol* [Internet]. 2018 Dec [cited 2022 Jan 18];84(12):2923–7. Available from: <https://onlinelibrary.wiley.com/doi/10.1111/bcp.13582>

12. Omar A, Ellen R, Sorisky A. Metformin-Associated Lactic Acidosis in a Patient with Normal Renal Function. *Canadian Journal of Diabetes* [Internet]. 2016 Aug [cited 2022 Jan 18];40(4):280-1. Available from: <https://linkinghub.elsevier.com/retrieve/pii/S1499267115300290>
13. Krowl L, Al-Khalisy H, Kaul P. Metformin-Induced Lactic Acidosis (MILA): Review of current diagnostic paradigm. *The American Journal of Emergency Medicine* [Internet]. 2018 May [cited 2022 Jan 18];36(5):908.e3-908.e5. Available from: <https://linkinghub.elsevier.com/retrieve/pii/S0735675718301116>
14. Kinoshita H, Yanai M, Ariyoshi K, Ando M, Tamura R. A patient with metformin-associated lactic acidosis successfully treated with continuous renal replacement therapy: a case report. *J Med Case Reports* [Internet]. 2019 Dec [cited 2022 Jan 18];13(1):371. Available from: <https://jmedicalcasereports.biomedcentral.com/articles/10.1186/s13256-019-2311-5>
15. Ryu S, Oh S-K, Son S-H, Jeong W-J, You Y-H, Ham Y-R. Reversible Acute Blindness in Suspected Metformin-Associated Lactic Acidosis. *The Journal of Emergency Medicine* [Internet]. 2019 Nov [cited 2022 Jan 18];57(5):e153-6. Available from: <https://linkinghub.elsevier.com/retrieve/pii/S0736467919305657>
16. Schwetz V, Eisner F, Schilcher G, Eller K, Plank J, Lind A, et al. Combined metformin-associated lactic acidosis and euglycemic ketoacidosis. *Wien Klin Wochenschr* [Internet]. 2017 Sep [cited 2022 Jan 18];129(17-18):646-9. Available from: <http://link.springer.com/10.1007/s00508-017-1251-6>
17. Gao J, Gu Z, Xu Y, Na Y. Peritoneal dialysis treatment of metformin-associated lactic acidosis in a diabetic nephropathy patient. *CN* [Internet]. 2016 Nov 1 [cited 2022 Jan 18];86(11):279-82. Available from: http://www.dustri.com/article_response_page.html?artId=14681&doi=10.5414/CN108859&L=0

18. Joseph CMC. Symptomatic Hypoglycemia During Treatment with a Therapeutic Dose of Metformin. *Am J Case Rep* [Internet]. 2021 Apr 28 [cited 2022 Jan 18];22. Available from: <https://www.amjcaserep.com/abstract/index/idArt/931311>
19. Pasquel FJ, Hinedi Z, Umpierrez GE, Klein R, Adigweme A, Coralli R, et al. Metformin-Associated Lactic Acidosis. *The American Journal of the Medical Sciences* [Internet]. 2015 Mar [cited 2022 Jan 18];349(3):263–7. Available from: <https://linkinghub.elsevier.com/retrieve/pii/S0002962915300999>
20. Ebrahim I, Blockman M. Metabolic acidosis in a patient with metformin overdose. *S Afr Med J* [Internet]. 2017 Jan 30 [cited 2022 Jan 18];107(2):110. Available from: <http://www.samj.org.za/index.php/samj/article/view/11809>
21. Sharma MP, Kar SK. Surreptitious metformin abuse in anorexia nervosa presenting as periodic hypoglycaemia. *Aust N Z J Psychiatry* [Internet]. 2015 Sep [cited 2022 Jan 18];49(9):851–2. Available from: <http://journals.sagepub.com/doi/10.1177/0004867415584642>
22. Graham RE, Cartner M, Winearls J. A severe case of vasoplegic shock following metformin overdose successfully treated with methylene blue as a last line therapy. *BMJ Case Reports* [Internet]. 2015 Jul 6 [cited 2022 Jan 18];bcr2015210229. Available from: <https://casereports.bmj.com/lookup/doi/10.1136/bcr-2015-210229>
23. Regolisti G, Antoniotti R, Fani F, Greco P, Fiaccadori E. Treatment of Metformin Intoxication Complicated by Lactic Acidosis and Acute Kidney Injury: The Role of Prolonged Intermittent Hemodialysis. *American Journal of Kidney Diseases* [Internet]. 2017 Aug [cited 2022 Jan 18];70(2):290–6. Available from: <https://linkinghub.elsevier.com/retrieve/pii/S0272638617300732>
24. Schuenemeyer JW, Jette-Kelly LA. A 35-Year-Old Woman With Shock, Pulseless Electrical Activity Arrest, and Hemodynamic Collapse. *Chest* [Internet]. 2019 Mar [cited 2022 Jan 18];155(3):e75–7. Available from: <https://linkinghub.elsevier.com/retrieve/pii/S0012369218322451>

25. Harding SA, Biary R, Hoffman RS, Su MK, Howland MA. A Pharmacokinetic Analysis of Hemodialysis for Metformin-Associated Lactic Acidosis. *J Med Toxicol* [Internet]. 2021 Jan [cited 2022 Jan 18];17(1):70–4. Available from: <https://link.springer.com/10.1007/s13181-020-00802-742>.
26. Hevesy MR. Metformin-Associated Lactic Acidosis: An Atypical Presentation. *Advanced Emergency Nursing Journal* [Internet]. 2017 Jan [cited 2022 Jan 18];39(1):26–30. Available from: <https://journals.lww.com/01261775-201701000-00005>
27. Zhang QC, Hastings C, Johnson K, Slaven E. Metformin-Associated Lactic Acidosis Presenting Like Acute Mesenteric Ischemia. *The Journal of Emergency Medicine* [Internet]. 2019 Nov [cited 2022 Jan 18];57(5):720–2. Available from: <https://linkinghub.elsevier.com/retrieve/pii/S073646791930335X>
28. Nakamura A, Suzuki K, Imai H, Katayama N. Metformin-associated lactic acidosis treated with continuous renal replacement therapy. *BMJ Case Reports* [Internet]. 2017 Feb 10 [cited 2022 Jan 18];bcr2016218318. Available from: <https://casereports.bmj.com/lookup/doi/10.1136/bcr-2016-218318>
29. Naunton M, Kyle G, Owoka F, Naunton-Boom K. Pharmacist review prevents evolving metformin-associated lactic acidosis. *J Clin Pharm Ther* [Internet]. 2014 Oct [cited 2022 Jan 18];39(5):567–70. Available from: <https://onlinelibrary.wiley.com/doi/10.1111/jcpt.12187>
30. Avcı D, Çetinkaya A, Karahan S, Oğuzhan N, Karagöz H, Başak M, et al. Suicide commitment with metformin: our experience with five cases. *Renal Failure* [Internet]. 2013 Jul [cited 2022 Jan 18];35(6):863–5. Available from: <http://www.tandfonline.com/doi/full/10.3109/0886022X.2013.801299>
31. Kopec KT, Kowalski MJ. Metformin-Associated Lactic Acidosis (MALA): Case Files of the Einstein Medical Center Medical Toxicology Fellowship. *J Med Toxicol* [Internet]. 2013 Mar [cited 2022 Jan 18];9(1):61–6. Available from: <http://link.springer.com/10.1007/s13181-012-0278-3>

32. Tay S, Lee I-L. Survival after cardiopulmonary arrest with extreme hyperkalaemia and hypothermia in a patient with metformin-associated lactic acidosis. *Case Reports* [Internet]. 2012 Dec 20 [cited 2022 Jan 18];2012(dec20 1):bcr2012007804–bcr2012007804. Available from: <https://casereports.bmj.com/lookup/doi/10.1136/bcr-2012-007804>
33. Cantrell FL, Nelson CL, Gary RD, McIntyre IM. Fatal Metformin Intoxication with Markedly Elevated Blood and Liver Concentrations. *Journal of Analytical Toxicology* [Internet]. 2012 Nov 1 [cited 2022 Jan 18];36(9):657–9. Available from: <http://academic.oup.com/jat/article/36/9/657/784873/Fatal-Metformin-Intoxication-with-Markedly>
34. Kalantar-Zadeh K, Uppot RN, Lewandrowski KB. Case 23-2013: A 54-Year-Old Woman with Abdominal Pain, Vomiting, and Confusion. Cabot RC, Rosenberg ES, Harris NL, Shepard J-AO, Cort AM, Ebeling SH, et al., editors. *N Engl J Med* [Internet]. 2013 Jul 25 [cited 2022 Jan 18];369(4):374–82. Available from: <http://www.nejm.org/doi/10.1056/NEJMcp120815459>.
35. Laforest C, Saint-Marcoux F, Amiel J-B, Pichon N, Merle L. Monitoring of metformin-induced lactic acidosis in a diabetic patient with acute kidney failure and effect of hemodialysis. *CP* [Internet]. 2013 Feb 1 [cited 2022 Jan 18];51(02):147–51. Available from: http://www.dustri.com/article_response_page.html?artId=10146&doi=10.5414/CP201728&L=0
36. Plumb B, Parker A, Wong P. Feeling blue with metformin-associated lactic acidosis. *Case Reports* [Internet]. 2013 Mar 1 [cited 2022 Jan 18];2013(mar01 1):bcr2013008855–bcr2013008855. Available from: <https://casereports.bmj.com/lookup/doi/10.1136/bcr-2013-008855>
37. St-Onge M, Ajmo I, Poirier D, Laliberté M. l-Carnitine for the Treatment of a Calcium Channel Blocker and Metformin Poisoning. *J Med Toxicol* [Internet]. 2013 Sep [cited 2022 Jan 18];9(3):266–9. Available from: <http://link.springer.com/10.1007/s13181-013-0301-3>

38. Makówka A, Zawiasa A, Nowicki M. Prescription-medication sharing among family members: an unrecognized cause of a serious drug adverse event in a patient with impaired renal function. *CN* [Internet]. 2015 Mar 1 [cited 2022 Jan 18];83 (2015)(03):196–200. Available from: http://www.dustri.com/article_response_page.html?artId=11336&doi=10.5414/CN108052&L=0
39. Weisberg LS. Lactic Acidosis in a Patient with Type 2 Diabetes Mellitus. *CJASN* [Internet]. 2015 Aug 7 [cited 2022 Jan 18];10(8):1476–83. Available from: <https://cjasn.asnjournals.org/lookup/doi/10.2215/CJN.10871014>
40. White S, Driver BE, Cole JB. Metformin-Associated Lactic Acidosis Presenting as Acute ST-Elevation Myocardial Infarction. *The Journal of Emergency Medicine* [Internet]. 2016 Jan [cited 2022 Jan 18];50(1):32–6. Available from: <https://linkinghub.elsevier.com/retrieve/pii/S0736467915010604>
41. Bolleku E, Idrizi A, Barbullushi M, Bajrami V, Likaj E, Thereska N, et al. Metformin-induced Lactic Acidosis Associated with Multiorgan Failure. *Med Arh* [Internet]. 2013 [cited 2022 Jan 18];67(5):381. Available from: <http://www.scopemed.org/fulltextpdf.php?mno=46182>
42. Bonsignore A, Pozzi F, Fraternali Orcioni G, Ventura F, Palmiere C. Fatal metformin overdose: case report and postmortem biochemistry contribution. *Int J Legal Med* [Internet]. 2014 May [cited 2022 Jan 18];128(3):483–92. Available from: <http://link.springer.com/10.1007/s00414-013-0927-3>
43. Gonçalves BM, Coelho D. Acidosis láctica asociada a metformina: un caso clínico que reporta una complicación grave en el período perioperatorio. *Revista Española de Anestesiología y Reanimación* [Internet]. 2019 Nov [cited 2022 Jan 18];66(9):483–6. Available from: <https://linkinghub.elsevier.com/retrieve/pii/S0034935619300696>
44. Morales A, Danziger J. Management Consideration in Drug-Induced Lactic Acidosis. *CJASN* [Internet]. 2020 Oct 7 [cited 2022 Jan 18];15(10):1511–2.

Available

from:

<https://cjasn.asnjournals.org/lookup/doi/10.2215/CJN.14781219>

45. Hooda A, Mehta A, Hannallah F. Metformin-associated lactic acidosis precipitated by liraglutide use: adverse effects of aggressive antihyperglycaemic therapy. *BMJ Case Rep* [Internet]. 2018 Nov [cited 2022 Jan 18];11(1):e227102. Available from: <https://casereports.bmj.com/lookup/doi/10.1136/bcr-2018-227102>
46. Ozeki T, Kawato R, Watanabe M, Minatoguchi S, Murai Y, Ryuge A, et al. A Fatal Case of Metformin-associated Lactic Acidosis. *Intern Med* [Internet]. 2016 [cited 2022 Jan 18];55(7):775–8. Available from: https://www.jstage.jst.go.jp/article/internalmedicine/55/7/55_55.5415/article
47. Pallavi R, Chaudhari A. Prolonged Hemodialysis: An Antidote for Metformin-Associated Lactic Acidosis. *American Journal of Therapeutics* [Internet]. 2016 Mar [cited 2022 Jan 18];23(2):e606–8. Available from: <https://journals.lww.com/00045391-201603000-00040>
48. Zibar L, Zibar K. Hemodialysis-refractory metformin-associated lactate acidosis with hypoglycemia, hypothermia, and bradycardia in a diabetic patient with belated diagnosis and chronic kidney disease. *CP* [Internet]. 2017 Apr 1 [cited 2022 Jan 18];55(04):348–51. Available from: https://www.dustri.com/article_response_page.html?artId=15173&doi=10.5414/CP202686&L=0
49. Margiani C, Zorcolo L, Mura P, Saba M, Restivo A, Scintu F. Metformin-associated lactic acidosis and temporary ileostomy: a case report. *J Med Case Reports* [Internet]. 2014 Dec [cited 2022 Jan 18];8(1):449. Available from: <https://jmedicalcasereports.biomedcentral.com/articles/10.1186/1752-1947-8-449>
50. Zheng L. Metformin as a Rare Cause of Drug-Induced Liver Injury, a Case Report and Literature Review. *American Journal of Therapeutics* [Internet]. 2016

- Jan [cited 2022 Jan 18];23(1):e315-7. Available from: <https://journals.lww.com/00045391-201601000-00052>
51. Altun E, Kaya B, Paydaş S, Sarıakçalı B, Karayaylalı İ. Lactic acidosis induced by metformin in a chronic hemodialysis patient with diabetes mellitus type 2: Metformin-induced lactic acidosis. *Hemodial Int* [Internet]. 2014 Apr [cited 2022 Jan 18];18(2):529-31. Available from: <https://onlinelibrary.wiley.com/doi/10.1111/hdi.12109>
52. Ansari RiyazuddinSH, Mady A, Qutub H, Althomaly E, Alzayer Z, Moulana AmnaAR. Metformin-related acidosis in a woman while performing Haj: A conservative approach. *Saudi J Kidney Dis Transpl* [Internet]. 2015 [cited 2022 Jan 18];26(1):125. Available from: <http://www.sjkdt.org/text.asp?2015/26/1/125/148759>
53. Galiero F, Consani G, Biancofiore G, Ruschi S, Forfori F. Metformin intoxication: Vasopressin's key role in the management of severe lactic acidosis. *The American Journal of Emergency Medicine* [Internet]. 2018 Feb [cited 2022 Jan 18];36(2):341.e5-341.e6. Available from: <https://linkinghub.elsevier.com/retrieve/pii/S0735675717308859>
54. Hussain MI, Hall BM, Depczynski B, Connor SJ. Acute renal failure and metformin-associated lactic acidosis following colonoscopy. *Diabetes Research and Clinical Practice* [Internet]. 2014 Jul [cited 2022 Jan 18];105(1):e6-8. Available from: <https://linkinghub.elsevier.com/retrieve/pii/S0168822714000163>
55. Liu S, Xu L, Ma J, Huang R, Lin T, Li Z, et al. High-volume continuous venovenous hemodiafiltration plus resin hemoperfusion improves severe metformin-associated toxicity. *J Diabetes Investig* [Internet]. 2018 Jul [cited 2022 Jan 18];9(4):975-8. Available from: <https://onlinelibrary.wiley.com/doi/10.1111/jdi.12757>
56. Ncomanzi D, Sicut RMR, Sundararajan K. Metformin-associated lactic acidosis presenting as an ischemic gut in a patient who then survived a cardiac arrest: a case report. *J Med Case Reports* [Internet]. 2014 Dec [cited 2022 Jan 18];8(1):159.

Available from:
<https://jmedicalcasereports.biomedcentral.com/articles/10.1186/1752-1947-8-159>

57. Tung ML, Tan LK. Long term use of metformin leading to vitamin B 12 deficiency. *Diabetes Research and Clinical Practice* [Internet]. 2014 Jun [cited 2022 Jan 18];104(3):e75–6. Available from: <https://linkinghub.elsevier.com/retrieve/pii/S0168822714000151>
58. Yamada T, Shojima N, Yamauchi T, Kadowaki T. Lactic acidosis due to attempted suicide with metformin overdose: A case report. *Diabetes & Metabolism* [Internet]. 2016 Sep [cited 2022 Jan 18];42(4):290–1. Available from: <https://linkinghub.elsevier.com/retrieve/pii/S1262363616303895>
59. Acquistapace G, Rossi M, Garbi M, Cosci P, Canetta C, Manelli A, et al. Acute metformin intoxication: 2012 experience of Emergency Departement of Lodi, Italy. *Clinical Chemistry and Laboratory Medicine (CCLM)* [Internet]. 2014 Jan 1 [cited 2022 Jan 18];52(10). Available from: <https://www.degruyter.com/document/doi/10.1515/cclm-2014-0208/html>
60. Cheng H, Ko KT, Tzang R. Metformin - induced first - episode psychosis in patient with polycystic ovary syndrome using drospirenone. *Psychiatry Clin Neurosci* [Internet]. 2019 Apr [cited 2022 Jan 18];73(4):196–196. Available from: <https://onlinelibrary.wiley.com/doi/10.1111/pcn.12825>
61. Lalau JD, Azzoug ML, Kajbaf F, Briet C, Desailoud R. Metformin accumulation without hyperlactataemia and metformin-induced hyperlactataemia without metformin accumulation. *Diabetes & Metabolism* [Internet]. 2014 Jun [cited 2022 Jan 18];40(3):220–3. Available from: <https://linkinghub.elsevier.com/retrieve/pii/S1262363613002334>
62. Mohiuddin S, Almaleki N, Ashraf M, Hussein M. Metformin-associated lactic acidosis in a peritoneal dialysis patient. *Saudi J Kidney Dis Transpl* [Internet]. 2015 [cited 2022 Jan 18];26(2):325. Available from: <http://www.sjkdt.org/text.asp?2015/26/2/325/152498>

63. Cigarran S, Rodriguez ML, Pousa M, Menendez H, Mendez MJ. Transient vision loss in a patient with severe metformin-associated lactic acidosis. QJM [Internet]. 2012 Aug 1 [cited 2022 Jan 18];105(8):781-3. Available from: <https://academic.oup.com/qjmed/article-lookup/doi/10.1093/qjmed/hcr109>
64. Kwok WC, Chan TC, Luk J, Chan F. Metformin - Associated Lactic Acidosis in an Older Adult After Colonoscopy: An Uncommon Trigger for a Rare Complication. J Am Geriatr Soc [Internet]. 2013 Dec [cited 2022 Jan 18];61(12):2257-8. Available from: <https://onlinelibrary.wiley.com/doi/10.1111/jgs.12585>
65. Seo JH, Lee DY, Hong CW, Lee IH, Ahn KS, Kang GW. Severe Lactic Acidosis and Acute Pancreatitis Associated with Cimetidine in a Patient with Type 2 Diabetes Mellitus Taking Metformin. Intern Med [Internet]. 2013 [cited 2022 Jan 18];52(19):2245-8. Available from: https://www.jstage.jst.go.jp/article/internalmedicine/52/19/52_52.0697/_article
66. Mujtaba M, Geara AS, Madhrira M, Agarwala R, Anderson H, Cheng J-T, et al. Toxicokinetics of metformin-associated lactic acidosis with continuous renal replacement therapy. Eur J Drug Metab Pharmacokinet [Internet]. 2012 Dec [cited 2022 Jan 18];37(4):249-53. Available from: <http://link.springer.com/10.1007/s13318-012-0104-y>
67. K. Miller D, J. Brinson A, Catalano G, C. Catalano M. Lactic Acidosis, Hypotension, and Sensorineural Hearing Loss Following Intentional Metformin Overdose. CDS [Internet]. 2011 Nov 1 [cited 2022 Jan 18];6(5):346-9. Available from: <http://www.eurekaselect.com/openurl/content.php?genre=article&issn=1574-8863&volume=6&issue=5&spage=346>
68. Correia CS, Bronander KA. Metformin-associated Lactic Acidosis Masquerading as Ischemic Bowel. The American Journal of Medicine [Internet]. 2012 May

[cited 2022 Jan 18];125(5):e9. Available from:
<https://linkinghub.elsevier.com/retrieve/pii/S0002934311009958>

69. Devetzis V, Passadakis P, Panagoutsos S, Theodoridis M, Thodis E, Georgoulidou A, et al. Metformin-related lactic acidosis in patients with acute kidney injury. *Int Urol Nephrol* [Internet]. 2011 Dec [cited 2022 Jan 18];43(4):1243–8. Available from: <http://link.springer.com/10.1007/s11255-010-9845-1>
70. Kelliher L, Fawcett W. Lactic acidosis in a diabetic patient treated with metformin undergoing hepatic resection. *British Journal of Hospital Medicine* [Internet]. 2012 Apr [cited 2022 Jan 18];73(4):232–3. Available from: <http://www.magonlinelibrary.com/doi/10.12968/hmed.2012.73.4.232>
71. van Sloten TT, Pijpers E, Stehouwer CDA, Brouwers MCGJ. Metformin-associated lactic acidosis in a patient with normal kidney function. *Diabetes Research and Clinical Practice* [Internet]. 2012 Jun [cited 2022 Jan 18];96(3):e57–8. Available from: <https://linkinghub.elsevier.com/retrieve/pii/S0168822711003950>
72. Mustafa E, Lai L, Lien Y-HH. Rapid Recovery from Acute Kidney Injury in a Patient with Metformin-associated Lactic Acidosis and Hypothermia. *The American Journal of Medicine* [Internet]. 2012 Feb [cited 2022 Jan 18];125(2):e1–2. Available from: <https://linkinghub.elsevier.com/retrieve/pii/S0002934311005481>
73. Hashmi T. Probable hepatotoxicity associated with the use of metformin in type 2 diabetes. *Case Reports* [Internet]. 2011 Sep 13 [cited 2022 Jan 18];2011(sep13 1):bcr0420114092–bcr0420114092. Available from: <https://casereports.bmj.com/lookup/doi/10.1136/bcr.04.2011.4092>
74. Akoglu H, Akan B, Piskinpaşa S, Karaca O, Dede F, Erdem D, et al. Metformin-associated lactic acidosis treated with prolonged hemodialysis. *The American Journal of Emergency Medicine* [Internet]. 2011 Jun [cited 2022 Jan

18];29(5):575.e3-575.e5. Available from:
<https://linkinghub.elsevier.com/retrieve/pii/S0735675710002494>

75. Aperis G, Paliouras C, Zervos A, Arvanitis A, Alivanis P. LACTIC ACIDOSIS AFTER CONCOMITANT TREATMENT WITH METFORMIN AND TENOFOVIR IN A PATIENT WITH HIV INFECTION: LACTIC ACIDOSIS AFTER CONCOMITANT TREATMENT WITH METFORMIN AND TENOFOVIR IN A PATIENT WITH HIV INFECTION. *Journal of Renal Care* [Internet]. 2011 Mar [cited 2022 Jan 18];37(1):25-9. Available from: <https://onlinelibrary.wiley.com/doi/10.1111/j.1755-6686.2011.00209.x>

76. Cone CJ, Bachyrycz AM, Murata GH. Hepatotoxicity Associated with Metformin Therapy in Treatment of Type 2 Diabetes Mellitus with Nonalcoholic Fatty Liver Disease. *Ann Pharmacother* [Internet]. 2010 Oct [cited 2022 Jan 18];44(10):1655-9. Available from: <http://journals.sagepub.com/doi/10.1345/aph.1P099>

77. Kreshak AA, Clark RF. Transient vision loss in a patient with metformin-associated lactic acidosis. *The American Journal of Emergency Medicine* [Internet]. 2010 Nov [cited 2022 Jan 18];28(9):1059.e5-1059.e7. Available from: <https://linkinghub.elsevier.com/retrieve/pii/S0735675710000264>

78. Dell'Aglio DM, Perino LJ, Todino JD, Algren DA, Morgan BW. Metformin Overdose With a Resultant Serum pH of 6.59: Survival Without Sequelae. *The Journal of Emergency Medicine* [Internet]. 2010 Jul [cited 2022 Jan 18];39(1):e77-80. Available from: <https://linkinghub.elsevier.com/retrieve/pii/S0736467907008529>

79. Giuliani E, Albertini G, Vaccari C, Barbieri A. pH 6.68--surviving severe metformin intoxication. *QJM* [Internet]. 2010 Nov 1 [cited 2022 Jan 18];103(11):887-90. Available from: <https://academic.oup.com/qjmed/article-lookup/doi/10.1093/qjmed/hcq049>

80. Rifkin SI, McFarren C, Juvvadi R, Weinstein SS. Prolonged Hemodialysis for Severe Metformin Intoxication. *Renal Failure* [Internet]. 2011 May [cited 2022

- Jan 18];33(4):459-61. Available from: <http://www.tandfonline.com/doi/full/10.3109/0886022X.2011.568132>
81. Arroyo AM, Walroth TA, Mowry JB, Kao LW. The MALAdy of Metformin Poisoning: Is CVVH the Cure? *American Journal of Therapeutics* [Internet]. 2010 Jan [cited 2022 Jan 18];17(1):96-100. Available from: <https://journals.lww.com/00045391-201001000-00021>
82. Carlon R, Dal Cero P, Corbanese U, Possamai C. Three cases of severe metformin-related lactic acidosis: *European Journal of Anaesthesiology* [Internet]. 2010 Jul [cited 2022 Jan 18];27(7):666-7. Available from: <http://journals.lww.com/00003643-201007000-00016>
83. Huberlant V, Laterre P-F, Hantson P. Nearly fatal metabolic acidosis: septic or toxic?: *European Journal of Emergency Medicine* [Internet]. 2010 Aug [cited 2022 Jan 18];17(4):243-4. Available from: <http://journals.lww.com/00063110-201008000-00015>
84. Saidi H, Mani M. Severe metabolic acidosis secondary to coadministration of creatine and metformin, a case report. *The American Journal of Emergency Medicine* [Internet]. 2010 Mar [cited 2022 Jan 18];28(3):388.e5-388.e6. Available from: <https://linkinghub.elsevier.com/retrieve/pii/S0735675709003854>
85. Lemyze M, Baudry JF, Collet F, Guinard N. Life threatening lactic acidosis. *BMJ* [Internet]. 2010 Mar 24 [cited 2022 Jan 18];340(mar24 2):c857-c857. Available from: <https://www.bmj.com/lookup/doi/10.1136/bmj.c857>
86. Akinci B, Yener S, Bengi G, Yesil S. Alterations of coagulation in metformin intoxication. *HJ* [Internet]. 2008 Oct 15 [cited 2022 Jan 18];7(4):325-9. Available from: <http://www.hormones.gr/310/article/alterations-of-coagulation-in-metformin-intoxication%E2%80%A6.html>
87. Brouwers MCGJ, Schaper N, Keeris L. Does glucose infusion exacerbate metformin-associated lactate acidosis? A case report. *Diabetes Research and Clinical Practice* [Internet]. 2009 Jul [cited 2022 Jan 18];85(1):e1-3. Available from: <https://linkinghub.elsevier.com/retrieve/pii/S0168822709001612>

88. Pan LTT, Maclaren G. Continuous Venovenous Haemodiafiltration for Metformin-induced Lactic Acidosis. *Anaesth Intensive Care* [Internet]. 2009 Sep [cited 2022 Jan 18];37(5):830-2. Available from: <http://journals.sagepub.com/doi/10.1177/0310057X0903700520>
89. Turkcuer I, Erdur B, Sari I, Yuksel A, Tura P, Yuksel S. Severe metformin intoxication treated with prolonged haemodialyses and plasma exchange: *European Journal of Emergency Medicine* [Internet]. 2009 Feb [cited 2022 Jan 18];16(1):11-3. Available from: <http://journals.lww.com/00063110-200902000-00003>
90. Yang P-W, Lin K-H, Lo S-H, Wang L-M, Lin H-D. Successful Treatment of Severe Lactic Acidosis Caused by a Suicide Attempt with a Metformin Overdose. *The Kaohsiung Journal of Medical Sciences* [Internet]. 2009 Feb [cited 2022 Jan 18];25(2):93-7. Available from: [http://doi.wiley.com/10.1016/S1607-551X\(09\)70047-6](http://doi.wiley.com/10.1016/S1607-551X(09)70047-6)
91. Chang L-C, Hung S-C, Yang C. The Case | A suicidal woman with delayed high anion gap metabolic acidosis. *Kidney International* [Internet]. 2009 Apr [cited 2022 Jan 18];75(7):757-8. Available from: <https://linkinghub.elsevier.com/retrieve/pii/S0085253815537708>
92. Fitzgerald E, Mathieu S, Ball A. Metformin associated lactic acidosis. *BMJ* [Internet]. 2009 Sep 16 [cited 2022 Jan 18];339(sep16 2):b3660-b3660. Available from: <https://www.bmj.com/lookup/doi/10.1136/bmj.b3660>
93. Jain V, Sharma D, Prabhakar H, Dash HH. Metformin-associated lactic acidosis following contrast media-induced nephrotoxicity: *European Journal of Anaesthesiology* [Internet]. 2008 Feb [cited 2022 Jan 18];25(2):166-7. Available from: <http://journals.lww.com/00003643-200802000-00013>
94. Jung EY, Cho HS, Seo JW, Kim DW, Kim H-J, Chang S-H, et al. Metformin-induced encephalopathy without lactic acidosis in a patient with contraindication for metformin. *Hemodialysis International* [Internet]. 2009 Apr

- [cited 2022 Jan 18];13(2):172-5. Available from: <https://onlinelibrary.wiley.com/doi/10.1111/j.1542-4758.2009.00358.x>
95. Papanas N, Monastiriotis C, Christakidis D, Maltezos E. METFORMIN AND LACTIC ACIDOSIS IN PATIENTS WITH TYPE 2 DIABETES - FROM PRIDE AND PREJUDICE TO SENSE AND SENSIBILITY. *Acta Clinica Belgica* [Internet]. 2009 Feb [cited 2022 Jan 18];64(1):42-8. Available from: <http://www.tandfonline.com/doi/full/10.1179/acb.2009.008>
96. Ashall V, Dawes T. Metformin and lactic acidosis. *British Journal of Anaesthesia* [Internet]. 2008 Dec [cited 2022 Jan 18];101(6):876-7. Available from: <https://linkinghub.elsevier.com/retrieve/pii/S0007091217341120>
97. Audia P, Feinfeld DA, Dubrow A, Winchester JF. Metformin-induced lactic acidosis and acute pancreatitis precipitated by diuretic, celecoxib, and candesartan-associated acute kidney dysfunction. *Clinical Toxicology* [Internet]. 2008 Jan [cited 2022 Jan 18];46(2):164-6. Available from: <http://www.tandfonline.com/doi/full/10.1080/15563650701355314>
98. Hong Y-C, O'Boyle CP, Chen I-C, Hsiao C-T, Kuan J-T. Metformin-Associated Lactic Acidosis in a Pregnant Patient. *Gynecol Obstet Invest* [Internet]. 2008 [cited 2022 Jan 18];66(2):138-41. Available from: <https://www.karger.com/Article/FullText/132463>
99. Kumar U, Chennavir B, Gopi A. Heeding clues to metformin-associated lactic acidosis: prompt response can save life. *Emergency Medicine Journal* [Internet]. 2008 Aug 1 [cited 2022 Jan 18];25(8):537-8. Available from: <https://emj.bmj.com/lookup/doi/10.1136/emj.2007.054213>
100. Prikis M, Mesler EL, Hood VL, Weise WJ. When a friend can become an enemy! Recognition and management of metformin-associated lactic acidosis. *Kidney International* [Internet]. 2007 Nov [cited 2022 Jan 18];72(9):1157-60. Available from: <https://linkinghub.elsevier.com/retrieve/pii/S0085253815527865>
101. EL-Hennawy AS, Jacob S, Mahmood AK. Metformin-Associated Lactic Acidosis Precipitated by Diarrhea. *American Journal of Therapeutics* [Internet]. 2007 Jul

- [cited 2022 Jan 18];14(4):403–5. Available from:
<https://journals.lww.com/00045391-200707000-00014>
102. Galea M, Jelacin N, Bramham K, White I. Severe lactic acidosis and rhabdomyolysis following metformin and ramipril overdose. *British Journal of Anaesthesia* [Internet]. 2007 Feb [cited 2022 Jan 18];98(2):213–5. Available from:
<https://linkinghub.elsevier.com/retrieve/pii/S0007091217355277>
103. Gambaro V, Dell'Acqua L, Farè F, Fidani M, Froidi R, Saligari E. A Case of Fatal Intoxication from Metformin. *J Forensic Sci* [Internet]. 2007 Jul [cited 2022 Jan 18];52(4):988–91. Available from:
<https://onlinelibrary.wiley.com/doi/10.1111/j.1556-4029.2007.00461.x>
104. Nyirenda MJ, Sandeep T, Grant I, Price G, McKnight JA. Severe acidosis in patients taking metformin-rapid reversal and survival despite high APACHE score. *Diabet Med* [Internet]. 2006 Apr [cited 2022 Jan 18];23(4):432–5. Available from: <https://onlinelibrary.wiley.com/doi/10.1111/j.1464-5491.2006.01813.x>
105. Alivanis P, Giannikouris I, Paliouras C, Arvanitis A, Volanaki M, Zervos A. Metformin-associated lactic acidosis treated with continuous renal replacement therapy. *Clinical Therapeutics* [Internet]. 2006 Mar [cited 2022 Jan 18];28(3):396–400. Available from:
<https://linkinghub.elsevier.com/retrieve/pii/S0149291806000695>
106. Guo PYF, Storsley LJ, Finkle SN. Dialysis Rounds A Dialysis Case Presentation and Discussion Edited by Roger A. Rodby: Severe Lactic Acidosis Treated with Prolonged Hemodialysis: Recovery After Massive Overdoses of Metformin: DIALYSIS IN MASSIVE METFORMIN OVERDOSES. *Seminars in Dialysis* [Internet]. 2006 Jan 17 [cited 2022 Jan 18];19(1):80–3. Available from:
<https://onlinelibrary.wiley.com/doi/10.1111/j.1525-139X.2006.00123.x>
107. Riesenman PJ, Braithwaite SS, Cairns BA. Metformin-Associated Lactic Acidosis in a Burn Patient: *Journal of Burn Care & Research* [Internet]. 2007 Mar [cited 2022 Jan 18];28(2):342–7. Available from:
<https://academic.oup.com/jbcr/article/28/2/342-347/4605528>

108. Tewari P, Nath S, Mazumdar G. Increased prothrombin time and lactic acidosis: Rofecoxib drug interaction with acenocoumarin and metformin. *Ann Card Anaesth* [Internet]. 2007 [cited 2022 Jan 18];10(1):58. Available from: <http://www.annals.in/text.asp?2007/10/1/58/37927>
109. Brassoe R, Elkmann T, Hempel M, Gravholt CH. Fulminant lactic acidosis in two patients with Type 2 diabetes treated with metformin. *Diabet Med* [Internet]. 2005 Oct [cited 2022 Jan 18];22(10):1451-3. Available from: <https://onlinelibrary.wiley.com/doi/10.1111/j.1464-5491.2005.01650.x>
110. Friesecke S, Abel P, Kraft M, Gerner A, Runge S. Combined renal replacement therapy for severe metformin-induced lactic acidosis. *Nephrology Dialysis Transplantation* [Internet]. 2006 Jul 1 [cited 2022 Jan 18];21(7):2038-9. Available from: <http://academic.oup.com/ndt/article/21/7/2038/1821773/Combined-renal-replacement-therapy-for-severe>
111. Schmidt A, Christensson A, Åkeson J. Intensive care treatment of severe mixed metabolic acidosis. *Acta Anaesthesiologica Scandinavica* [Internet]. 2005 Mar [cited 2022 Jan 18];49(3):411-4. Available from: <https://onlinelibrary.wiley.com/doi/10.1111/j.1399-6576.2005.00586.x>
112. Shenoy C. Metformin-Associated Lactic Acidosis Precipitated by Acute Renal Failure. *The American Journal of the Medical Sciences* [Internet]. 2006 Jan [cited 2022 Jan 18];331(1):55-7. Available from: <https://linkinghub.elsevier.com/retrieve/pii/S000296291532872X>
113. Bryant SM, Cumpston K, Lipsky MS, Patel N, Leikin JB. Metformin-Associated Respiratory Alkalosis: *American Journal of Therapeutics* [Internet]. 2004 May [cited 2022 Jan 18];11(3):236-7. Available from: <http://journals.lww.com/00045391-200405000-00014>
114. Panzer U, Kluge S, Kreymann G, Wolf G. Combination of intermittent haemodialysis and high-volume continuous haemofiltration for the treatment of severe metformin-induced lactic acidosis. *Nephrology Dialysis Transplantation*

- [Internet]. 2004 Aug 1 [cited 2022 Jan 18];19(8):2157-8. Available from: <https://academic.oup.com/ndt/article-lookup/doi/10.1093/ndt/gfh337>
115. Dawson D, Conlon C. Case Study: Metformin-Associated Lactic Acidosis. *Diabetes Care* [Internet]. 2003 Aug 1 [cited 2022 Jan 18];26(8):2471-2. Available from: <https://diabetesjournals.org/care/article/26/8/2471/22663/Case-Study-Metformin-Associated-Lactic>
116. Nammour FE, Fayad NF, Peikin SR. Metformin-Induced Cholestatic Hepatitis. *Endocrine Practice* [Internet]. 2003 Jul [cited 2022 Jan 18];9(4):307-9. Available from: <https://linkinghub.elsevier.com/retrieve/pii/S1530891X20402095>
117. von Mach M-A, Sauer O, Sacha Weilemann L. Experiences of a Poison Center with Metformin-Associated Lactic Acidosis. *Exp Clin Endocrinol Diabetes* [Internet]. 2004 May 4 [cited 2022 Jan 18];112(04):187-90. Available from: <http://www.thieme-connect.de/DOI/DOI?10.1055/s-2004-817931>
118. Edwards CMB, Barton MA, Snook J, David M, Mak VHF, Chowdhury TA. Metformin - associated lactic acidosis in a patient with liver disease. *QJM: An International Journal of Medicine* [Internet]. 2003 Apr [cited 2022 Jan 18];96(4):315-6. Available from: <https://academic.oup.com/qjmed/article-lookup/doi/10.1093/qjmed/hcg049>
119. Gjedde S, Christiansen A, Pedersen SB, Rungby J. Survival Following a Metformin Overdose of 63 g: A Case Report. *Pharmacology & Toxicology* [Internet]. 2003 Aug [cited 2022 Jan 18];93(2):98-9. Available from: <https://onlinelibrary.wiley.com/doi/10.1034/j.1600-0773.2003.930207.x>
120. Mallick S. Metformin induced acute pancreatitis precipitated by renal failure. *Postgraduate Medical Journal* [Internet]. 2004 Apr 1 [cited 2022 Jan 18];80(942):239-40. Available from: <https://pmj.bmj.com/lookup/doi/10.1136/pgmj.2003.011957>
121. Nisse P, Mathieu - Nolf M, Deveaux M, Forceville X, Combes A. A Fatal Case of Metformin Poisoning. *Journal of Toxicology: Clinical Toxicology* [Internet]. 2003

- Jan [cited 2022 Jan 18];41(7):1035-6. Available from: <http://www.tandfonline.com/doi/full/10.1081/CLT-120026533>
122. Schier JG, Hoffman RS, Nelson LS. Metformin-Induced Acidosis Due to a Warfarin Adverse Drug Event. *Ann Pharmacother* [Internet]. 2003 Jul [cited 2022 Jan 18];37(7-8):1145-1145. Available from: <http://journals.sagepub.com/doi/10.1345/aph.1D074>
123. Worth L, Elliott J, Anderson J, Sasadeusz J, Street A, Lewin S. A Cautionary Tale: Fatal Lactic Acidosis Complicating Nucleoside Analogue and Metformin Therapy. *Clinical Infectious Diseases* [Internet]. 2003 Jul 15 [cited 2022 Jan 18];37(2):315-6. Available from: <https://academic.oup.com/cid/article-lookup/doi/10.1086/375850>
124. Ahmad S. Recovery from pH 6.38: lactic acidosis complicated by hypothermia. *Emergency Medicine Journal* [Internet]. 2002 Mar 1 [cited 2022 Jan 18];19(2):169-71. Available from: <https://emj.bmj.com/lookup/doi/10.1136/emj.19.2.169>
125. Barrueto F, Meggs WJ, Barchman MJ. Clearance of Metformin by Hemofiltration in Overdose. *Journal of Toxicology: Clinical Toxicology* [Internet]. 2002 Jan [cited 2022 Jan 18];40(2):177-80. Available from: <http://www.tandfonline.com/doi/full/10.1081/CLT-120004407>
126. Chang C-T, Chen Y-C, Fang J-T, Huang C-C. HIGH ANION GAP METABOLIC ACIDOSIS IN SUICIDE: DON'T FORGET METFORMIN INTOXICATION—TWO PATIENTS' EXPERIENCES. *Renal Failure* [Internet]. 2002 Jan [cited 2022 Jan 18];24(5):671-5. Available from: <http://www.tandfonline.com/doi/full/10.1081/JDI-120013973>
127. Vander T, Hallevy H, Ifergane G, Herishanu YO. Metformin-induced encephalopathy without lactic acidosis. *Diabet Med* [Internet]. 2004 Feb [cited 2022 Jan 18];21(2):194-5. Available from: <http://doi.wiley.com/10.1046/j.1464-5491.2003.01087.x>
128. Zitzmann S, Reimann IR, Schmechel H. Severe hypoglycemia in an elderly patient treated with metformin. *CP* [Internet]. 2002 Mar 1 [cited 2022 Jan 18];40(03):108-

10. Available from:
http://www.dustri.com/article_response_page.html?artId=5712&doi=10.5414/CPP40108&L=0
129. Khan JK, Pallaki M, Tolbert SR, Hornick TR. Lactic Acidemia Associated with Metformin. *Ann Pharmacother* [Internet]. 2003 Jan [cited 2022 Jan 18];37(1):66-9. Available from: <http://journals.sagepub.com/doi/10.1345/aph.1C183>
130. Mears SC, Lipsett PA, Brager MD, Riley LH. Metformin-Associated Lactic Acidosis After Elective Cervical Spine Fusion: A Case Report. *Spine* [Internet]. 2002 Nov [cited 2022 Jan 18];27(22):E482-4. Available from: <http://journals.lww.com/00007632-200211150-00019>
131. Price G. Metformin lactic acidosis, acute renal failure and rofecoxib. *British Journal of Anaesthesia* [Internet]. 2003 Dec [cited 2022 Jan 18];91(6):909-10. Available from:
<https://linkinghub.elsevier.com/retrieve/pii/S0007091217362773>
132. Franzetti I, Paolo D, Marco G, Emanuela M, Elisabetta Z, Renato U. Possible synergistic effect of metformin and enalapril on the development of hyperkalemic lactic acidosis. *Diabetes Research and Clinical Practice* [Internet]. 1997 Dec [cited 2022 Jan 18];38(3):173-6. Available from:
<https://linkinghub.elsevier.com/retrieve/pii/S0168822797000983>
133. Kruse JA. Metformin-associated lactic acidosis. *The Journal of Emergency Medicine* [Internet]. 2001 Apr [cited 2022 Jan 18];20(3):267-72. Available from:
<https://linkinghub.elsevier.com/retrieve/pii/S0736467900003206>
134. Heaney D, Majid A, Junor B. Bicarbonate haemodialysis as a treatment of metformin overdose. *Nephrology Dialysis Transplantation* [Internet]. 1997 May 1 [cited 2022 Jan 18];12(5):1046-7. Available from:
<https://academic.oup.com/ndt/article-lookup/doi/10.1093/ndt/12.5.1046>
135. Teale KFH, Devine A, Stewart H, Harper NJH. The management of metformin overdose. *Anaesthesia* [Internet]. 1998 Jul [cited 2022 Jan 18];53(7):698-701.

Available from: <https://onlinelibrary.wiley.com/doi/abs/10.1046/j.1365-2044.1998.436-az0549.x>

136. Brady WJ, Carter CT. Metformin overdose. *The American Journal of Emergency Medicine* [Internet]. 1997 Jan [cited 2022 Jan 18];15(1):107–8. Available from: <https://linkinghub.elsevier.com/retrieve/pii/S0735675797900735>
137. Doorenbos CJ, Bosma RJ, Lamberts PJJ. Use of urea containing dialysate to avoid disequilibrium syndrome, enabling intensive dialysis treatment of a diabetic patient with renal failure and severe metformin induced lactic acidosis. *Nephrology Dialysis Transplantation* [Internet]. 2001 Jun 1 [cited 2022 Jan 18];16(6):1303–4. Available from: <http://academic.oup.com/ndt/article/16/6/1303/1889615>
138. Jurovich MR, Wooldridge JD, Force RW. Metformin-Associated Nonketotic Metabolic Acidosis. *Ann Pharmacother* [Internet]. 1997 Jan [cited 2022 Jan 18];31(1):53–5. Available from: <http://journals.sagepub.com/doi/10.1177/106002809703100108>
139. Lalau JD, Race J-M, Brinquin L. Lactic Acidosis in Metformin Therapy: Relationship between plasma metformin concentration and renal function. *Diabetes Care* [Internet]. 1998 Aug 1 [cited 2022 Jan 18];21(8):1366–7. Available from: <https://diabetesjournals.org/care/article/21/8/1366/20179/Lactic-Acidosis-in-Metformin-Therapy-Relationship>
140. Pepper GM, Schwartz M. Lactic Acidosis Associated With Glucophage Use in a Man With Normal Renal and Hepatic Function. *Diabetes Care* [Internet]. 1997 Feb 1 [cited 2022 Jan 18];20(2):232–3. Available from: <https://diabetesjournals.org/care/article/20/2/232/18847/Lactic-Acidosis-Associated-With-Glucophage-Use-in>
141. Chan N, Fauvel N, Feher M. Non-steroidal anti-inflammatory drugs and metformin: a cause for concern? *The Lancet* [Internet]. 1998 Jul [cited 2022 Jan 18];352(9123):201. Available from: <https://linkinghub.elsevier.com/retrieve/pii/S0140673605778065>

142. Darwich MS, Allaf W. Metabolic acidosis. Postgraduate Medical Journal [Internet]. 1998 Jul 1 [cited 2022 Jan 18];74(873):427-8. Available from: <https://pmj.bmj.com/lookup/doi/10.1136/pgmj.74.873.427>
143. Hulisz DT, Bonfiglio MF, Murray RD. Metformin-Associated Lactic Acidosis. The Journal of the American Board of Family Medicine [Internet]. 1998 May 1 [cited 2022 Jan 18];11(3):233-6. Available from: <http://www.jabfm.org/cgi/doi/10.3122/15572625-11-3-233>
144. Mercker SK, Maier C, Neumann G, Wulf H. Lactic Acidosis as a Serious Perioperative Complication of Antidiabetic Biguanide Medication with Metformin. Anesthesiology [Internet]. 1997 Oct 1 [cited 2022 Jan 18];87(4):1003-5. Available from: <https://pubs.asahq.org/anesthesiology/article/87/4/1003/36236/Lactic-Acidosis-as-a-Serious-Perioperative>
145. Al-Jebawi AF, Lassman MN, Abourizk NN. Lactic Acidosis With Therapeutic Metformin Blood Level in a Low-Risk Diabetic Patient. Diabetes Care [Internet]. 1998 Aug 1 [cited 2022 Jan 18];21(8):1364-5. Available from: <https://diabetesjournals.org/care/article/21/8/1364/20131/Lactic-Acidosis-With-Therapeutic-Metformin-Blood>
146. Babich MM, Pike I, Shiffman ML. Metformin-induced Acute Hepatitis. The American Journal of Medicine [Internet]. 1998 May [cited 2022 Jan 18];104(5):490-2. Available from: <https://linkinghub.elsevier.com/retrieve/pii/S0002934398000886>
147. Schmidt R, Horn E, Richards J, Stamatakis M. Survival after metformin-associated lactic acidosis in peritoneal dialysis-dependent renal failure. The American Journal of Medicine [Internet]. 1997 May [cited 2022 Jan 18];102(5):486-8. Available from: <https://linkinghub.elsevier.com/retrieve/pii/S0002934397894442>
148. Hermann LS, Magnusson S, Möller B, Casey C, Tucker GT, Woods HF. Lactic Acidosis during Metformin Treatment in an Elderly Diabetic Patient with

- Impaired Renal Function. *Acta Medica Scandinavica* [Internet]. 2009 Apr 24 [cited 2022 Jan 18];209(1-6):519-20. Available from: <https://onlinelibrary.wiley.com/doi/10.1111/j.0954-6820.1981.tb11639.x>
149. Pearlman BL, Fenves AZ, Emmett M. Metformin-associated lactic acidosis. *The American Journal of Medicine* [Internet]. 1996 Jul [cited 2022 Jan 18];101(1):109-10. Available from: <https://linkinghub.elsevier.com/retrieve/pii/S0002934397894223>
150. Safadi R, Dranitzki-Elhalel M, Popovtzer M, Ben-Yehuda A. Metformin-Induced Lactic Acidosis Associated with Acute Renal Failure. *Am J Nephrol* [Internet]. 1996 [cited 2022 Jan 18];16(6):520-2. Available from: <https://www.karger.com/Article/FullText/169052>
151. Gan SC. Biguanide-associated lactic acidosis. Case report and review of the literature. *Archives of Internal Medicine* [Internet]. 1992 Nov 1 [cited 2022 Jan 18];152(11):2333-6. Available from: <http://archinte.ama-assn.org/cgi/doi/10.1001/archinte.152.11.2333>
152. Khan IH, Catto GR, MacLeod AM. Severe lactic acidosis in patient receiving continuous ambulatory peritoneal dialysis. *BMJ* [Internet]. 1993 Oct 23 [cited 2022 Jan 18];307(6911):1056-7. Available from: <https://www.bmj.com/lookup/doi/10.1136/bmj.307.6911.1056>
153. Lalau JD, Westeel PF, Debussche X, Dkissi H, Tolani M, Coevoet B, et al. Bicarbonate haemodialysis: an adequate treatment for lactic acidosis in diabetics treated by metformin. *Intensive Care Med* [Internet]. 1987 [cited 2022 Jan 18];13(6). Available from: <http://link.springer.com/10.1007/BF00257680>
154. McLelland J. Recovery from Metformin Overdose. *Diabetic Medicine* [Internet]. 1985 Sep [cited 2022 Jan 18];2(5):410-1. Available from: <https://onlinelibrary.wiley.com/doi/10.1111/j.1464-5491.1985.tb00668.x>
155. Phillips PJ, Scicchitano R, Clarkson AR, Gilmore HR. Metformin Associated Lactic Acidosis. *Australian and New Zealand Journal of Medicine* [Internet]. 1978 Jun

- [cited 2022 Jan 18];8(3):281-4. Available from:
<https://onlinelibrary.wiley.com/doi/10.1111/j.1445-5994.1978.tb04524.x>
156. Assan R, Heuclin Ch, Ganeval D, Bismuth Ch, George J, Girard JR. Metformin-induced lactic acidosis in the presence of acute renal failure. *Diabetologia* [Internet]. 1977 May [cited 2022 Jan 18];13(3):211-7. Available from:
<http://link.springer.com/10.1007/BF01219702>
157. Tymms DJ, Leatherdale BA. Lactic acidosis due to metformin therapy in a low risk patient. *Postgraduate Medical Journal* [Internet]. 1988 Mar 1 [cited 2022 Jan 18];64(749):230-1. Available from:
<https://pmj.bmj.com/lookup/doi/10.1136/pgmj.64.749.230>
158. Günay E, Kalkan Z, Selim Ay M, Yuksel E, Berekatoglu N, Yaman M. Successful Dialysis Treatment of Metformin Associated Lactic Acidosis, Report of Two Cases. *Iran J Kidney Dis*. 2020 Mar;14(2):160-4.
159. Nestor C, Nasim S, Coyle N, Canavan C. Laboratory-Confirmed Metformin-Associated Lactic Acidosis. *Ir Med J*. 2019 Sep 12;112(8):992.
160. Ortiz-Brizuela E, Pérez-Patrigéon S, Recillas-Gispert C, Gómez-Pérez FJ. Lactic Acidosis Complicating Metformin and Non-Nucleoside Reverse Transcriptase Inhibitor Combination Therapy: A Smoldering Threat in the Post-HAART Era. *Rev Invest Clin*. 2015 Aug;67(4):273-4.
161. Sehra S, Jaggi S, Sehra D, Aggarwal R, Saraswat V, Juneja D. Management of Sitagliptin and Metformin Combination Toxic Overdose. *J Assoc Physicians India*. 2016 Nov;64(11):80-1.
162. Sertbas M, Sertbas Y, Ordu O, Berber E, Ozen B, Ozdemir A. Myocardial injury and acute renal failure associated with lactic acidosis due to suicide attempt with metformin. *J Pak Med Assoc*. 2016 Feb;66(2):223-5.
163. Girasole F, Piccolo G, Timpanelli R, Calanna M, Piazza F, Vecchio S. AKI with serious state of acidosis in diabetic patients treated with metformin. *G Ital Nefrol*. 2013 Oct;30(5):gin/30.5.6.

164. Tsitsios T, Sotirakopoulos N, Armentzioiou K, Kalogiannidou I, Kolaggis A, Mavromatidis K. Metformin induced severe hypophosphatemia in a patient on hemodialysis. *Saudi J Kidney Dis Transpl.* 2010 Sep;21(5):923-6.
165. Al-Makadma YS, Riad T. Successful management of high-dose metformin intoxication. Role of vasopressin in the management of severe lactic acidosis. *Middle East J Anaesthesiol.* 2010 Oct;20(6):873-5.
166. Bruijstens LA, van Luin M, Buscher-Jungerhans PMM, Bosch FH. Reality of severe metformin-induced lactic acidosis in the absence of chronic renal impairment. *Neth J Med.* 2008 May;66(5):185-90.
167. Di Grande A, Vancheri F, Giustolisi V, Giuffrida C, Narbone G, Licata M, et al. Metformin-induced lactic acidosis in a type 2 diabetic patient with acute renal failure. *Clin Ter.* 2008 Apr;159(2):87-9.
168. Talwalkar PG, Deotale P. Severe hypoglycemia in a patient with type 2 diabetes mellitus on metformin monotherapy. *J Assoc Physicians India.* 2007 Jun;55:458-9.
169. Fox GN. Metformin in type 2 diabetes. *J Fam Pract.* 2004 Nov;53(11):900.
170. Kutoh E. Possible metformin-induced hepatotoxicity. *Am J Geriatr Pharmacother.* 2005 Dec;3(4):270-3.
171. Chu C-K, Chang Y-T, Lee B-J, Hu S-Y, Hu W-H, Yang D-Y. Metformin-associated lactic acidosis and acute renal failure in a type 2 diabetic patient. *J Chin Med Assoc.* 2003 Aug;66(8):505-8.
172. Schure PJCM, de Gooijer A, van Zanten ARH. Unexpected survival from severe metformin-associated lactic acidosis. *Neth J Med.* 2003 Oct;61(10):331-3.
173. Beis SJ, Goshman LM, Newkirk GL. Risk factors for metformin-associated lactic acidosis. *WMJ.* 1999 Aug;98(4):56-7.
174. Gueriguian J, Green L, Misbin RI, Stadel B, Fleming GA. Efficacy of metformin in non-insulin-dependent diabetes mellitus. *N Engl J Med.* 1996 Jan 25;334(4):269; author reply 269-270.

175. Chalopin JM, Tanter Y, Besancenot JF, Cabanne JF, Rifle G. Treatment of metformin-associated lactic acidosis with closed recirculation bicarbonate-buffered hemodialysis. *Arch Intern Med*. 1984 Jan;144(1):203-5.
176. Hofkens PJ, De Winter S, Vanbrabant P. Metformin-associated lactic acidosis (MALA): case report. *Acta Clin Belg*. 2011 Oct;66(5):390-2.
177. Habimana-Jordana A, López-Corominas V, Barceló-Martín B, Gomila-Muñiz I, Martínez-Sánchez L. Metabolic lactic acidosis as a sign of voluntary poisoning in adolescents. *Anales de Pediatría (English Edition)* [Internet]. 2019 Feb 1;90(2):121-3. Available from: <https://www.sciencedirect.com/science/article/pii/S2341287918302242>
178. Ives Tallman C, Zhang Y, Black N, Lynch K, Fayed M, Armenian P. Refractory vasodilatory shock secondary to metformin overdose supported with VA ECMO. *Toxicology Reports* [Internet]. 2022 Jan 1;9:64-7. Available from: <https://www.sciencedirect.com/science/article/pii/S2214750021002195>
179. Nannapaneni N, Elkhider A, Steinberg J. Hypoglycemic Hemiparesis: The Stroke Masquerader. *The American Journal of Medicine* [Internet]. 2014 Sep 1;127(9):e13. Available from: <https://www.sciencedirect.com/science/article/pii/S0002934314004677>
180. Ortiz-Lasa M, Gonzalez-Castro A, Peñasco Martín Y. Lactic acidosis associated (or induced by) metformin. *Medicina Clínica (English Edition)* [Internet]. 2017 Nov 9;149(9):415-6. Available from: <https://www.sciencedirect.com/science/article/pii/S238702061730640X>
181. Pasquel FJ, Hinedi Z, Umpierrez GE, Klein R, Adigweme A, Coralli R, et al. Metformin-Associated Lactic Acidosis. *The American Journal of the Medical Sciences* [Internet]. 2015 Mar 1;349(3):263-7. Available from: <https://www.sciencedirect.com/science/article/pii/S0002962915300999>
182. Shenoy C. Metformin-Associated Lactic Acidosis Precipitated by Acute Renal Failure. *The American Journal of the Medical Sciences* [Internet]. 2006 Jan

1;331(1):55-7.

Available

from:

<https://www.sciencedirect.com/science/article/pii/S000296291532872X>

183. McCabe DJ, Baker S, Stellpflug SJ. Hemodialysis in metformin-associated lactic acidosis due to acute overdose in a metformin-naïve patient. *The American Journal of Emergency Medicine* [Internet]. 2018 Sep 1;36(9):1721.e1-1721.e2.

Available

from:

<https://www.sciencedirect.com/science/article/pii/S0735675718304522>

184. Price G. Metformin lactic acidosis, acute renal failure and rofecoxib. *British Journal of Anaesthesia* [Internet]. 2003 Dec 1;91(6):909-10. Available from:

<https://www.sciencedirect.com/science/article/pii/S0007091217362773>