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Digitalis Historical Background and Current Status

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Abstract: The earliest recorded treatment of digitalis is typically credited to William Witherings investigation on the foxglove, which was published in 1785. The rich background of digitalis is intriguing. Yet, there is evidence of some awareness of herbs used for remedies for heart failure with complications that have effects similar to those of digitalis dating back to roman times. In relation to this, the foxglove's natural components (Digitalis purpurea and Digitalis lanata) also include ouabain, a quick-acting glycoside typically derived from Strophanthus gratus. These substances are known as cardiotonic steroids. These medications are effective sodium-potassium adenosine triphosphatase antagonists. Digitalis as well as its metabolites, specifically digoxin, served as the gold standard of treatment for CHF during the duration of the 20th century. As the century came to a close, however, several concerns—particularly those related to ensuring enhanced safety—were raised regarding their usage as additional therapies for CHF, such as lowering the left ventricle's preload. An important medication used to treat cardiac arrhythmias and high blood pressure is still digitalis glycosides. The properties of the all cardioactive glycosides enhance the myocardial fibre contractile strength in a manner that is similar. Almost 100% of digoxin is absorbed, has a T^{4} life of 5 to 7 days, and is mainly excreted in the urine as cardioinactive metabolites with just 8% of it being converted to digoxin. The inhibition of membrane Na^+/K^+ ATPase and its resulting impacts on calcium movement are thought to be the reason behind this. Digitalis and certain medications can interact, most commonly with diuretics that cause hypokalaemia or hypomagnesaemia. The management of cardiac arrhythmias following digitalis toxicity is mainly possible by favourable interactions with antiarrhythmic pharmaceuticals (lignocaine, phenytoin), however the efficiency of other medications, such propranolol, is occasionally constrained by their adverse inotropic effects.

Keywords: Digitalis, digitalis pharmacokinetics, Pharmacological actions, digitalis adulterants, digitalis toxicities

REFERENCES

- [1]. Withering W. An account of the foxglove, and some of its medical uses. Cambridge University Press; 2014 Sep 25.
- [2]. Schoner W, Scheiner-Bobis G. Endogenous and exogenous cardiac glycosides and their mechanisms of action. American journal of cardiovascular drugs. 2007 May; 7:173-89.
- [3]. de Wit M, Schoemaker NJ. Clinical approach to avian cardiac disease. InSeminars in Avian and Exotic Pet Medicine 2005 Jan 1 (Vol. 14, No. 1, pp. 6-13). WB Saunders.
- [4]. Taboulet P, Baud FJ, Bismuth C. Clinical features and management of digitalis poisoning—rationale for immunotherapy. Journal of Toxicology: Clinical Toxicology. 1993 Jan 1;31(2):247-60.
- [5]. Lip GY, Metcalfe MJ, Dunn FG. Diagnosis and treatment of digoxin toxicity. Postgraduate medical journal. 1993 May;69(811):337.

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- **[6].** Arold G, Donath F, Maurer A, Diefenbach K, Bauer S, Henneicke-von Zepelin HH, Friede M, Roots I. No relevant interaction with alprazolam, caffeine, tolbutamide, and digoxin by treatment with a low-hyperform St John's wort extract. Planta medica. 2005 Apr;71(04):331-7
- [7]. Marcus FI, Kapadia GJ, Kapadia GG. The metabolism of digoxin in normal subjects. Journal of Pharmacology and Experimental Therapeutics. 1964 Aug 1;145(2):203-9.
- [8]. Rathore SS, Curtis JP, Wang Y, Bristow MR, Krumholz HM. Association of serum digoxin concentration and outcomes in patients with heart failure. Jama. 2003 Feb 19;289(7):871-8.
- [9]. Patel S. Plant-derived cardiac glycosides: Role in heart ailments and cancer management. Biomedicine & Pharmacotherapy. 2016 Dec 1;84:1036-41.
- **[10].** Perne A, Muellner MK, Steinrueck M, Craig-Mueller N, Mayerhofer J, Schwarzinger I, Sloane M, Uras IZ, Hoermann G, Nijman SM, Mayerhofer M. Cardiac glycosides induce cell death in human cells by inhibiting general protein synthesis. PloS one. 2009 Dec 16;4(12):e8292.
- [11]. Prassas I, Diamandis EP. Novel therapeutic applications of cardiac glycosides. Nature reviews Drug discovery. 2008 Nov;7(11):926-35.
- [12]. Stenkvist BJ. Is digitalis a therapy for breast carcinoma?. Oncology reports. 1999 May 1;6(3):493-9.
- [13]. Manunta P, Ferrandi M, Bianchi G, Hamlyn JM. Endogenous ouabain in cardiovascular function and disease. Journal of hypertension. 2009 Jan 1;27(1):9-18.
- [14]. Hansen O. No evidence for a role in signal-transduction of Na+/K+-ATPase interaction with putative endogenous ouabain. European journal of biochemistry. 2003 May;270(9):1916-9.
- [15]. Bagrov AY, Shapiro JI, Fedorova OV. Endogenous cardiotonic steroids: physiology, pharmacology, and novel therapeutic targets. Pharmacological reviews. 2009 Mar 1;61(1):9-38.
- [16]. Cho J, Lee YJ, Kim JH, Kim SI, Kim SS, Choi BS, Choi JH. Antiviral activity of digoxin and ouabain against SARS-CoV-2 infection and its implication for COVID-19. Scientific reports. 2020 Oct 1;10(1):16200.
- [17]. Ouadid H, Albat B, Nargeot J. Calcium currents in diseased human cardiac cells. Journal of cardiovascular pharmacology. 1995 Feb 1;25(2):282-91.
- [18]. Manunta P, Ferrandi M, Bianchi G, Hamlyn JM. Endogenous ouabain in cardiovascular function and disease. Journal of hypertension. 2009 Jan 1;27(1):9-18.
- [19]. Best LS, Bierzychudek P. Pollinator foraging on foxglove (Digitalis purpurea): a test of a new model. Evolution. 1982 Jan 1:70-9.
- [20]. Wang H, Haas M, Liang M, Cai T, Tian J, Li S, Xie Z. Ouabain assembles signaling cascades through the caveolar Na+/K+-ATPase. Journal of Biological Chemistry. 2004 Apr 23;279(17):17250-9.
- [21]. Buckalew VM. Role of endogenous digitalis-like factors in the clinical manifestations of severe preeclampsia: a systematic review. Clinical Science. 2018 Jun 21;132(12):1215-42.
- [22]. Hixson-Wallace J. Digoxin toxicity: a review. US Pharm. 2006;2:28-36.
- [23]. Whayne TF. Clinical use of digitalis: a state of the art review. American journal of cardiovascular drugs. 2018 Dec;18:427-40.\
- [24]. Kanji S, MacLean RD. Cardiac glycoside toxicity: more than 200 years and counting. Critical care clinics. 2012 Oct 1;28(4):527-35.
- [25]. Goldman P. Herbal medicines today and the roots of modern pharmacology. Annals of internal medicine. 2001 Oct 16;135(8_Part_1):594-600.
- [26]. Ehle M, Patel C, Giugliano RP. Digoxin: clinical highlights: a review of digoxin and its use in contemporary medicine. Critical pathways in cardiology. 2011 Jun 1;10(2):93-8.
- [27]. Perloff MD, Von Moltke LL, Störmer E, Shader RI, Greenblatt DJ. Saint John's wort: an in vitro analysis of P-glycoprotein induction due to extended exposure. British journal of pharmacology. 2001 Dec;134(8):1601-8.
- [28]. Bilia AR, Gallori S, Vincieri FF. St. John's wort and depression: efficacy, safety and tolerability-an update. Life sciences. 2002 May 17;70(26):3077-96.

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- [29]. e Vries M, Seppala LJ, Daams JG, van de Glind EM, Masud T, van der Velde N, Blain H, Bousquet J, Bucht G, Caballero-Mora MA, van der Cammen T. Fall-risk-increasing drugs: a systematic review and metaanalysis: I. Cardiovascular drugs. Journal of the American Medical Directors Association. 2018 Apr 1;19(4):371-e1.
- [30]. Duan R, Du W, Guo W. EZH2: a novel target for cancer treatment. Journal of hematology & oncology. 2020 Dec;13(1):1-2.
- [31]. Osman MH, Farrag E, Selim M, Osman MS, Hasanine A, Selim A. Cardiac glycosides use and the risk and mortality of cancer; systematic review and meta-analysis of observational studies. PloS one. 2017 Jun 7;12(6):e0178611.
- [32]. Kepp O, Menger L, Vacchelli E, Adjemian S, Martins I, Ma Y, Sukkurwala AQ, Michaud M, Galluzzi L, Zitvogel L, Kroemer G. Anticancer activity of cardiac glycosides: At the frontier between cell-autonomous and immunological effects. Oncoimmunology. 2012 Dec 1;1(9):1640-2.

