


Antithrombotics in concept maps: an innovative approach to Pharmacology education in Dentistry

Gustavo Lucas Ribeiro Batista¹

 [0000-0002-1841-9809](https://orcid.org/0000-0002-1841-9809)

Rafael Lara Brasil¹

 [0000-0002-5118-4829](https://orcid.org/0000-0002-5118-4829)

Erica Negrini Lia¹

 [0000-0001-5691-415X](https://orcid.org/0000-0001-5691-415X)

¹Departamento de Odontologia, Faculdade de Ciências da Saúde, Universidade de Brasília (UnB), Brasília, Distrito Federal, Brasil.

Correspondence:

Gustavo Lucas Ribeiro Batista
E-mail: gustavolrb2008@gmail.com

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Abstract This study aimed to develop concept maps (CMs) on antithrombotic drugs for undergraduate dentistry teaching with information based on updated scientific evidence. Initially, consulting the scientific literature carried out a list of the fundamental concepts. Next, information on antithrombotic drugs was organized, and the focal questions were defined. The concepts were then inserted into the CMs, and the conceptual relationships were established. Then, searches were carried out in the PubMed, Scopus, and Embase databases for systematic reviews published in the last ten years using the terms “antithrombotics”, “antiplatelet”, “anticoagulants”, “antithrombotic therapy”, “dentistry” and “dental surgery”. In total, 23 articles were found, 12 were excluded, and 11 articles were included in the study. Subsequently, the CMs layouts were organized using the CmapTools software (v. 6.04). Eight CMs were developed on the hemostatic process, blood coagulation cascade, mechanism of action of anticoagulants and antiplatelet agents, and drug and food interactions. The CMs produced contain essential information for the training of Dentistry students on a topic of great importance, especially for surgical cases. Future studies will be conducted to test the effectiveness of these CMs on learning.

Descriptors: Education, Dental. Anticoagulants. Platelet Aggregation Inhibitors. Hemostasis.

Antitrombóticos en mapas conceptuales: un enfoque innovador para la enseñanza de Farmacología en Odontología

Resumen El objetivo de este estudio fue desarrollar mapas conceptuales (MCs) sobre fármacos antitrombóticos dirigidos a la enseñanza de la Odontología con información basada en evidencia científica actualizada. Inicialmente se identificaron los conceptos fundamentales consultando la literatura científica. A continuación, se organizó la información relacionada con los fármacos antitrombóticos y se definieron las preguntas focales. Los conceptos se insertaron en los MCs y se establecieron relaciones conceptuales. Para mejorar los conceptos se realizaron búsquedas en las bases de datos PubMed, Scopus y Embase de revisiones sistemáticas publicadas en los últimos diez años utilizando los términos *antithrombotics*, *antiplatelet*, *anticoagulants*, *antithrombotic therapy*, *dentistry* y *dental surgery*. En total se encontraron 23 artículos, 12 fueron excluidos y por tanto se incluyeron 11 artículos. Posteriormente se organizaron los diseños de MCs mediante la aplicación CmapTools (versión 6.04). Se desarrollaron ocho MCs sobre el proceso hemostático, la cascada de coagulación sanguínea, el mecanismo de acción de los anticoagulantes y antiplaquetarios y las interacciones entre medicamentos y alimentos. Los MCs producidos contienen información esencial para la formación de los estudiantes de Odontología en un tema de gran importancia, especialmente para los casos quirúrgicos. Se realizarán estudios futuros para probar la eficacia de los MCs en el aprendizaje.

Descriptor: Educación en Odontología. Anticoagulantes. Inhibidores de la Agregación Plaquetaria. Hemostasia.

Antitrombóticos em mapas conceituais: uma abordagem inovadora para o ensino de Farmacologia em Odontologia

Resumo O objetivo desse estudo foi desenvolver mapas conceituais (MCs) sobre fármacos antitrombóticos direcionados ao ensino de Odontologia com informações baseadas em evidências científicas atualizadas. Inicialmente os conceitos

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fundamentais foram identificados por meio de consulta à literatura científica. Em seguida, informações relacionadas aos fármacos antitrombóticos foram organizadas e as perguntas focais definidas. Os conceitos foram inseridos nos MCs e estabelecidas as relações conceituais. Para aprimorar os conceitos foram realizadas buscas nas bases de dados PubMed, Scopus e Embase por revisões sistemáticas publicadas nos últimos dez anos utilizando os termos *antithrombotics antiplatelet anticoagulants antithrombotic therapy dentistry e dental surgery*. Ao total foram encontrados 23 artigos. 12 foram excluídos e, portanto, 11 artigos foram incluídos. Subsequentemente, os *layouts* dos MCs foram organizados por meio do aplicativo CmapTools (versão 6.04). Foram desenvolvidos oito MCs sobre processo hemostático, cascata da coagulação sanguínea, mecanismo de ação dos anticoagulantes e antiagregantes plaquetários, interação medicamentosa e alimentar. Os MCs produzidos contém informações essenciais para a formação do aluno de Odontologia acerca de um tema de grande importância, em especial para casos cirúrgicos. Estudos futuros serão conduzidos para testar a eficácia dos MCs sobre o aprendizado.

Descritores: Educação em Odontologia. Anticoagulantes. Inibidores da Agregação Plaquetária. Hemostasia.

INTRODUCTION

There are topics of great interest in Dentistry, such as anticoagulants and antiplatelet agents, considering the increasing prescription for patients who have had or are at risk of thromboembolic events¹. However, a considerable portion of dentists still do not understand the mechanism of action of these drugs and fear uncontrollable bleeding during and after dental surgical procedures, thus avoiding immediate patient care¹.

Given their complexity and difficulty of understanding, concept maps (CMs) can be used as a strategy for their teaching²⁻⁴. CMs are graphic organizers representing knowledge through propositions, composed of three essential elements: the initial and final concepts, and the linking term (LT)². The use of CMs has significantly benefited the health sciences field due to their ability to make connections between concepts which increases critical thinking, reflection, and the student's ability to fully practice metalearning, which means "learning to learn"⁴.

There are few studies on using CMs in teaching Pharmacology, especially for dental students. Given the above, this study aimed to develop CMs on antithrombotic drugs at undergraduate Dentistry education, with information based on updated scientific evidence.

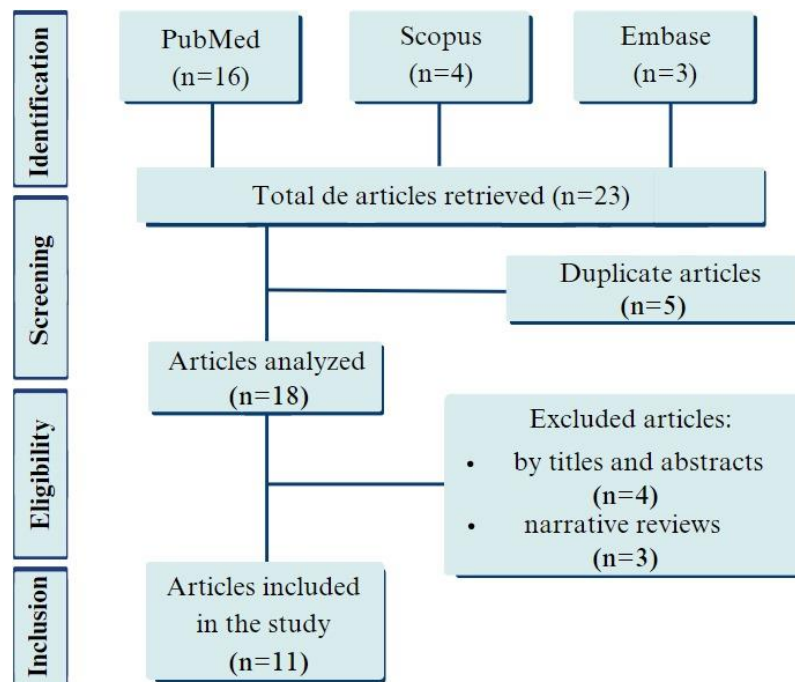
METHODS

Concept maps (CMs) were developed on the pharmacodynamics of drugs involving anticoagulant and antiplatelet therapies, drug and food interactions, and their implications in dental practice.

To this end, a list of fundamental concepts regarding the basic and clinical Pharmacology of anticoagulants and antiplatelet agents was initially compiled using book chapters^{5,6}. Next, the focal questions (FQs) were defined, concepts were inserted into the map, and conceptual relationships were established. Finally, the ideas were reviewed, and the concepts were refined through literature searches in the PubMed, Scopus, and Embase databases for systematic reviews published in the last ten years (Table 1). A total of 23 articles were found, with 12 excluded (five duplicates, four unrelated to the topic, and three narrative reviews). In total, 11 articles were included in the study⁷⁻¹⁷ (Figure 1). Subsequently, the Propositional Clarity Table (PCT)² was used, and the map layout was organized.

Table 1. Search strategies used for PubMed, Scopus, and Embase databases.

Database	Search on 01/11/2022
PubMed	(((((antithrombotics) OR (antithrombotic therapy)) OR (antiplatelet)) OR (anticoagulants)) AND (dentistry)) AND (dental surgery) Filters: ten last years and systematic revisions
Scopus	(TITLE-ABS-KEY (antithrombotic OR "antithrombotic therapy" OR antiplatelet OR anticoagulants AND dentistry AND "dental surgery") AND TITLE-ABS-KEY ("systematic review")) AND PUBYEAR > 2012 AND PUBYEAR > 2012
Embase	(antithrombotics OR 'antithrombotic therapy' OR antiplatelet OR anticoagulants) AND dentistry AND 'dental surgery' AND (2013:py OR 2014:py OR 2016:py OR 2017:py OR 2018:py OR 2019:py OR 2020:py OR 2021:py) AND 'systematic review'/de

**Figure 1.** Study selection flowchart.

RESULTS

The first FQ generated the first CM (Figure 2), which reviews the coagulation cascade's main points. The second FQ generated the second CM, which addresses the mechanisms of action of different anticoagulants (Figure 3). The third FQ supported the construction of the third CM on the hemostatic process (Figure 4). The fourth FQ generated the fourth CM on the mechanisms of action of different antiplatelet agents (Figure 5). The drug and food interactions of direct oral anticoagulants (DOACs), antiplatelet agents, Vitamin K antagonists, and heparins were presented in the remaining CMs (Figures 5, 6, 7, and 8, respectively).

The CMs should be read starting from the proposition formed by two concepts (initial and final) connected by an arrow, with a linking term (LT) in the middle, indicating the direction of the reading. It allows for defining the initial concept, the conceptual relationship, and the final concept of the proposition. Example: COAGULATION CASCADE-----is divided into---->INTRINSIC PATHWAY.

The first CM (Figure 2) presents the blood coagulation cascade, showing all the steps of the three pathways: intrinsic (in green), extrinsic (in blue), and common (in yellow). The extrinsic pathway is responsible for most of the coagulation *in vivo*, while the intrinsic pathway is mainly *in vitro*, and both converge at the activation point of factor X. Factor Xa and Thrombin (IIa) are the linking points to the second CM (Figure 3).

The second concept map (CM) (Figure 3) shows the mechanisms of action of the principal anticoagulants and their

sites of action, Thrombin (IIa), Antithrombin III, Factor Xa, and Vitamin K, in the coagulation cascade. The linking points between the first and second CMs are Factor Xa and Thrombin (IIa).

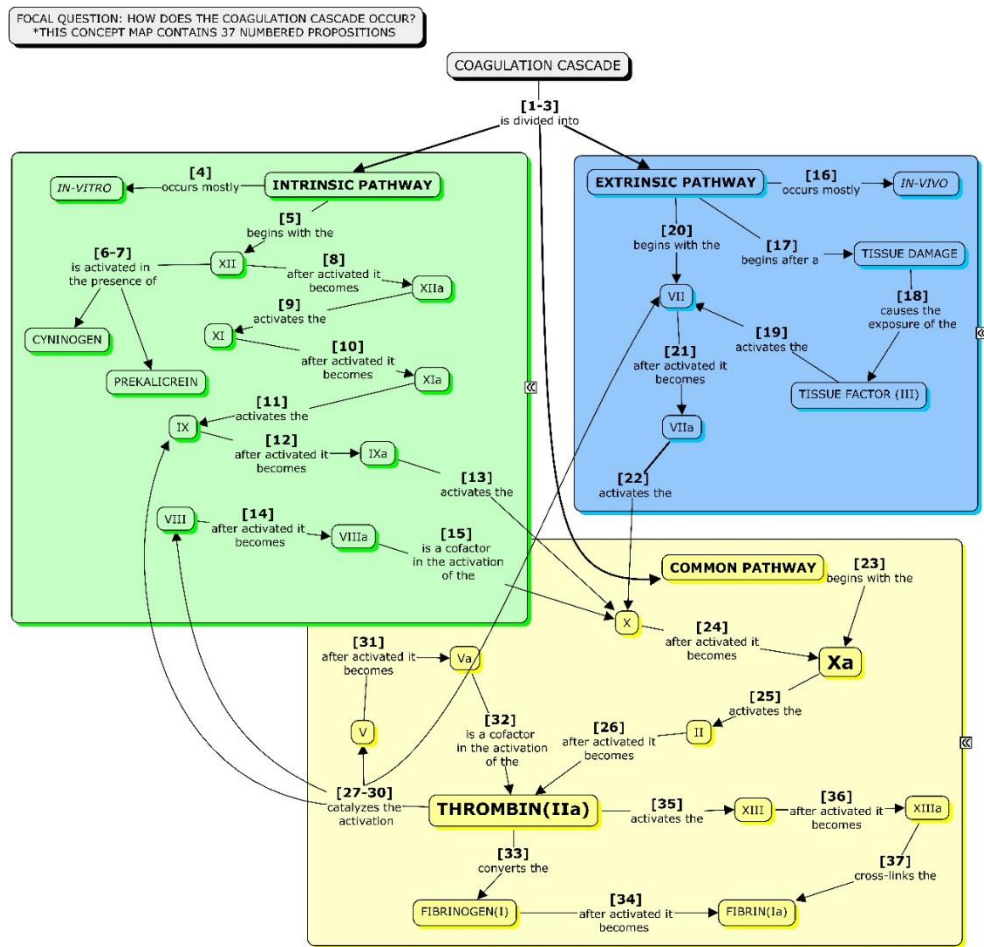


Figure 2. Concept Map of the Blood Coagulation Cascade. The concepts refer to blood coagulation factors, with the term “factor” omitted (e.g., factor X). The term “a” following the factors means activated (e.g., factor Xa, read as activated factor X).

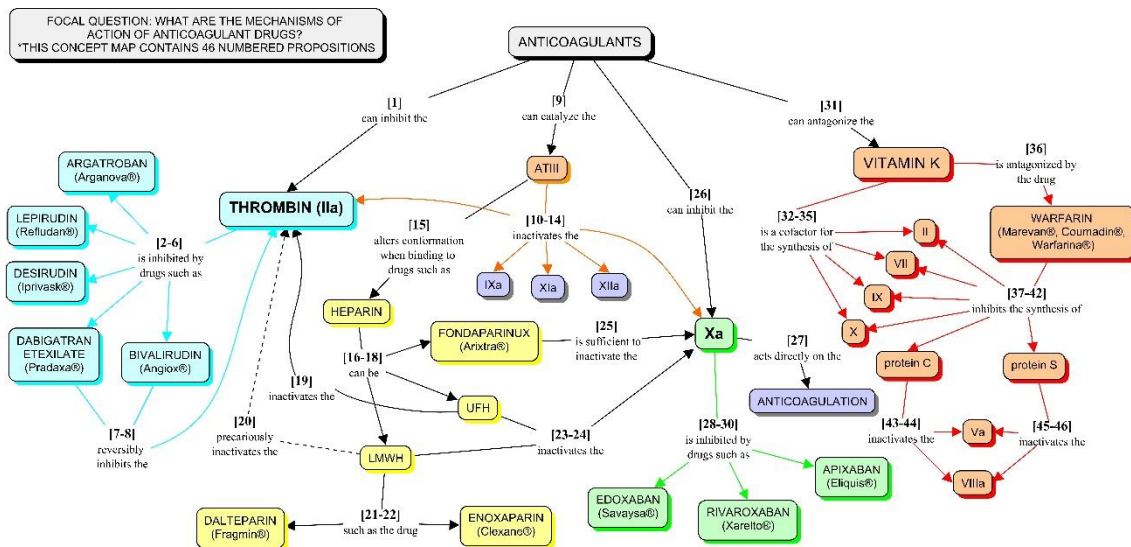


Figure 3. Concept Map of the mechanisms of action of Anticoagulants. Thrombin (IIa) inhibitors are in blue; Heparins in yellow; Factor Xa inhibitors in green; Vitamin K antagonists in red; Antithrombin III (ATIII) inhibitors in orange; and other factors and the concept of anticoagulation in purple. LMWH (Low Molecular Weight Heparin), UFH (Unfractionated Heparin).

The third CM (Figure 4) structures the hemostatic process divided into four phases: vasoconstriction (in red), primary hemostasis (in yellow), secondary hemostasis (in blue), and tertiary hemostasis (in green). This division is merely didactic, as these phases can occur simultaneously. The concepts ADP, TxA2, and Platelet Aggregation link points to the fourth CM (Figure 5).

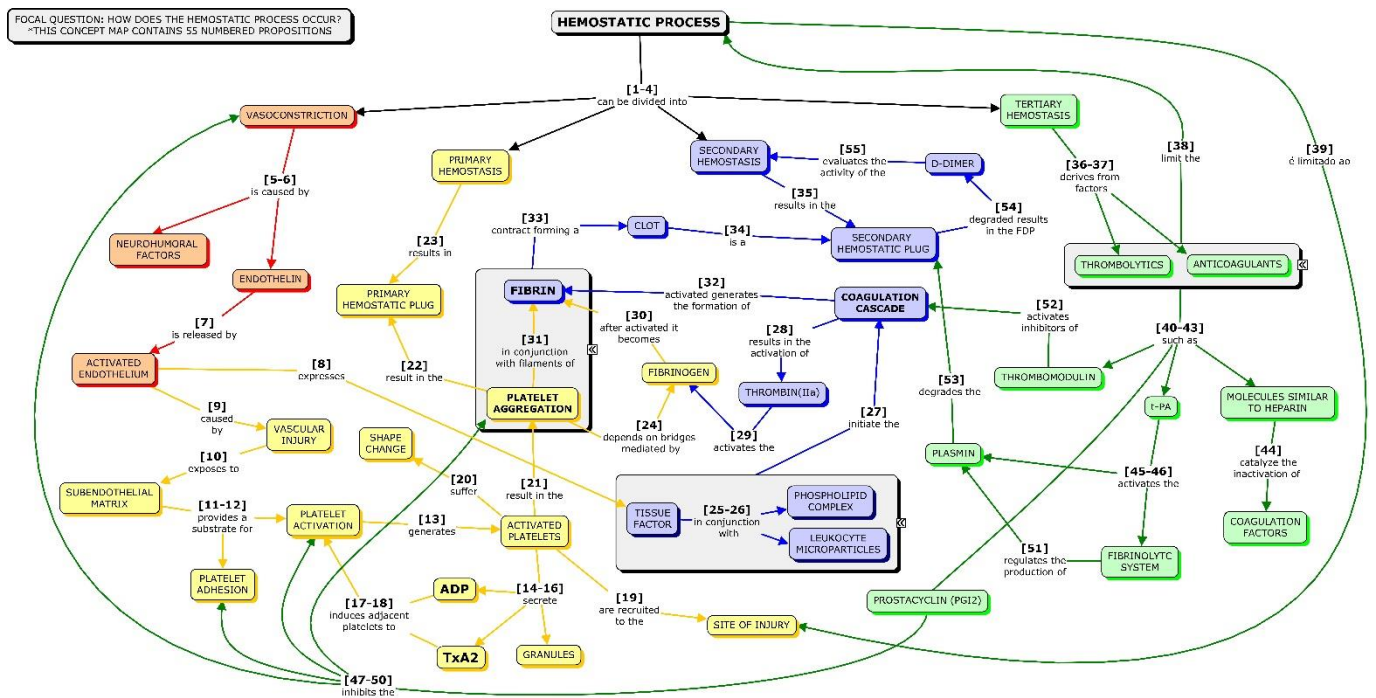


Figure 4. Concept Map of the Hemostatic Process Scheme. ADP (Adenosine Diphosphate), TxA2 (Thromboxane A2), t-PA (tissue plasminogen activator), and FDP (Fibrin Degradation Product).

The fourth concept map (CM) (Figure 5) illustrates the mechanisms of action of the main antiplatelet agents and their sites of action, such as COX1, TxA2; GPIIb/IIIa receptors, platelet aggregation via fibrinogen bridges; and ADP receptors (P2Y12), platelet activation. The fourth CM links to the third CM (Figure 4) through the COX1, ADP receptors, and GPIIb/IIIa receptors (concepts).

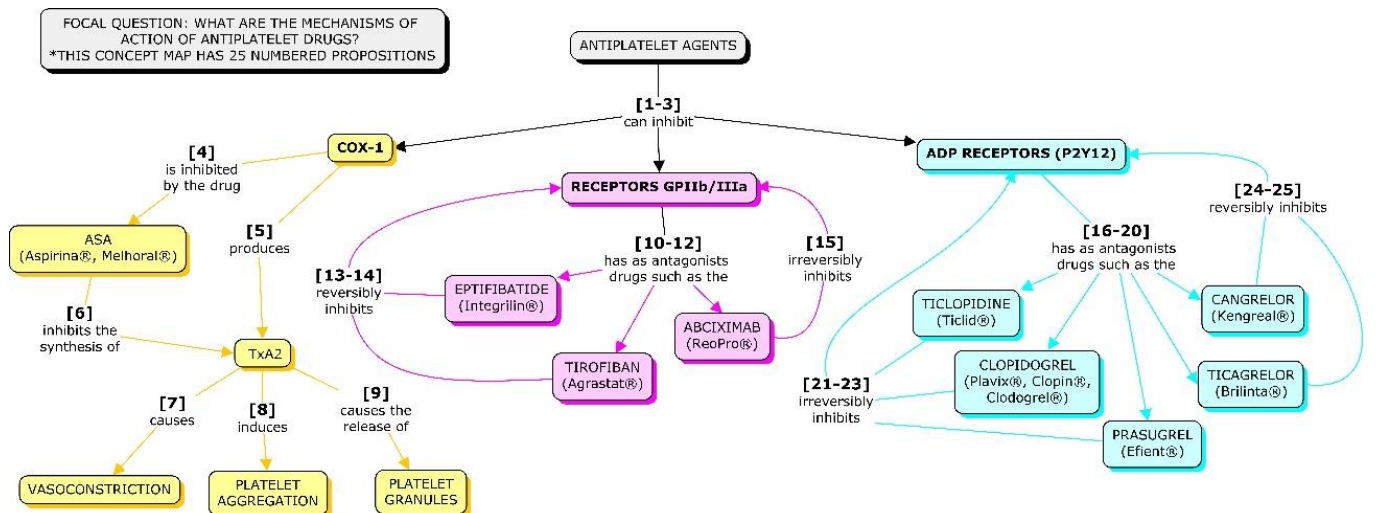


Figure 5. Concept Map of the mechanisms of action of Antiplatelet Agents. COX-1 (cyclooxygenase-1), ASA (acetylsalicylic acid) and TxA2 (thromboxane A2), ADP (Adenosine Diphosphate), GPIIb/IIIa Receptors (Glycoprotein IIb/IIIa Receptors), ADP Receptors – P2Y12 (Adenosine Diphosphate Receptors - P2Y12). Colors were used to sectorize the key points of the CM, which are the concepts COX1 (in yellow), GPIIb/IIIa Receptors (in pink), and ADP Receptors – P2Y12 (in blue).

CMs numbered 5 to 8 (Figures 6 to 9) highlight the principal drug and food interactions of anticoagulants and antiplatelet agents such as antibiotics, azole antifungals, analgesics, macrolides, anticonvulsants, nonsteroidal anti-inflammatory drugs (NSAIDs), barbiturates, foods rich in vitamin K, and alcohol. Additionally, in these CMs, the intensity of the interaction is presented by colors, such as lower, moderate, or higher. The intensity of the interaction was established using the academic license database of DrugBank, provided free of charge by the University of Alberta¹⁸. The concepts in CMs numbered 5 to 8 were not numbered as they only present examples of drugs, making a reading sequence unnecessary. The color intensity indicates the degree of interaction intensity. In shades of yellow (lower), orange (moderate), and dark orange (higher), drugs and foods that increase the effect of the drugs in each CM and increase the risk of bleeding are presented; in shades of light green (lower), green (moderate), and dark green (higher), drugs and foods that decrease the effect of the drugs in each CM are presented.

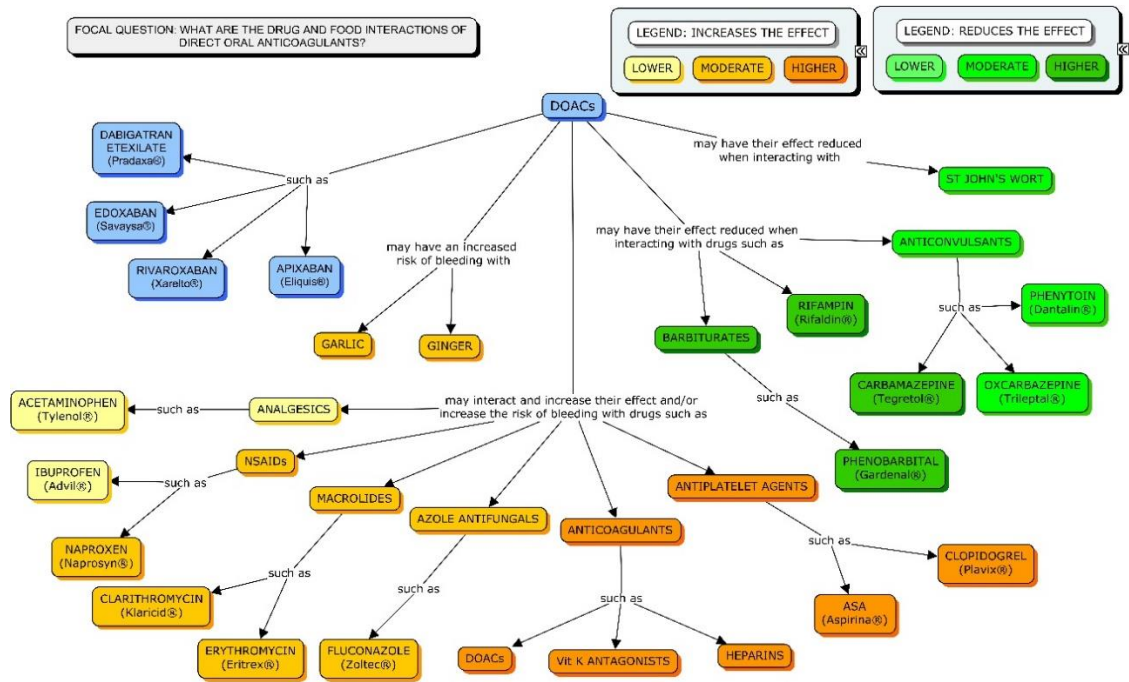


Figure 6. Concept Map of drug and food interactions of DOACs (Direct Oral Anticoagulants), NSAIDs (Nonsteroidal Anti-inflammatory Drugs). Examples of DOACs are highlighted in blue.

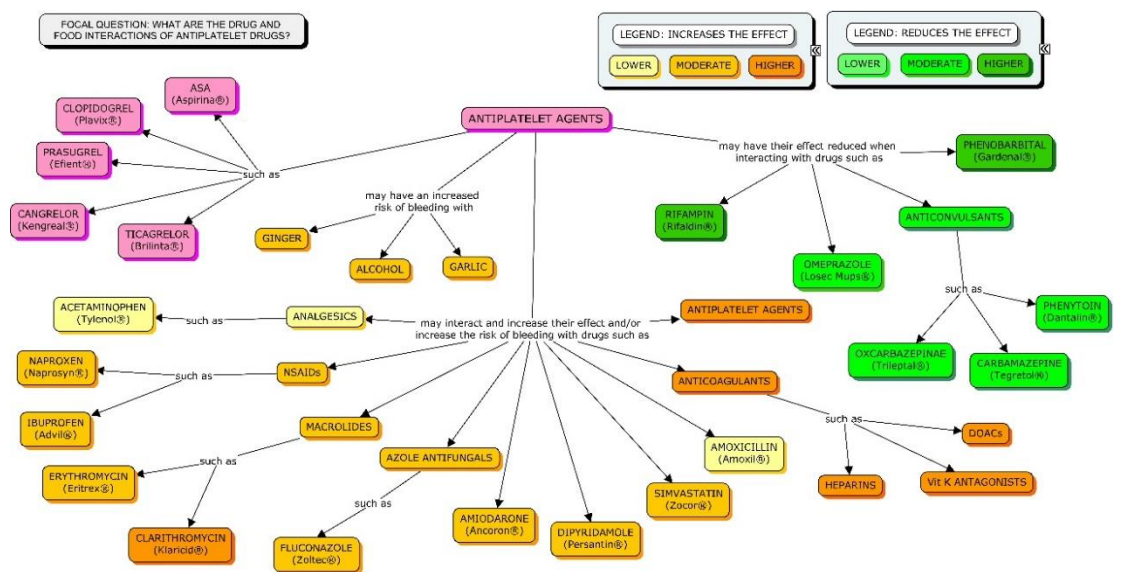


Figure 7. Concept Map of drug and food interactions of Antiplatelet Agents. ASA (Acetylsalicylic Acid), DOACs (Direct Oral Anticoagulants), NSAIDs (Nonsteroidal Anti-inflammatory Drugs). Examples of Antiplatelet Agents are highlighted in pink.

possible drug and food interactions with anticoagulants and antiplatelet agents. This information is essential for dental practice, as it allows for safe and informed decision-making when dealing with patients who use these medications.

It is important to emphasize that concept maps differ from mind maps, because they include concepts (initial and final) and LTs. The concept consists of one or more words that define regularities perceived in objects and events². The LT explains the conceptual relationship between the initial and final concepts, ideally using a conjugated verb to define the relationship². However, specifically regarding the CMs constructed in the study, it was decided to insert examples of drugs at the end of the propositions, which justifies the absence of verbs in these LTs.

The use of CMs as a teaching tool has been widely studied, given its effectiveness in promoting meaningful learning^{3,20}. By visualizing the relationships between concepts, students can better understand the organization and structure of knowledge²⁰, identify gaps in their understanding, and integrate new information more efficiently².

The CMs present hierarchically distributed concepts, with more general concepts at the beginning (top) and more specific ones at the end (bottom)^{2,21}. It is necessary to define a FQ^{2,21} to delimit the subject to be addressed by the CM, which will help evaluate whether all concepts fit the theme to limit the scope of the CM, thus avoiding overly extensive CMs. The FQ can be compared to the title of an essay. In a didactic manner, and as a guide for the reader, the propositions were numbered, and colors were inserted to sectorize groups of concepts that integrate different categories.

Considering the educational objectives of knowledge organization, study, and review, the produced Concept Maps (CMs) were based on the most recurrent applications of CMs in the literature, such as inserting new content or themes, organizing information hierarchically, describing complex processes, reviewing content from the most important concepts, synthesizing extensive content, addressing doubts and providing timely feedback². Therefore, the student can use the already constructed CM as support material during their studies and reviews. It is important to emphasize that CMs are configured as instructional materials, as they are produced through systematic consultations in specialized literature. In our study, the construction of CMs will allow them to be available to students and professionals to stimulate initial learning. Once students receive the ready CM, they will perform its dynamic reading and, thus, can dedicate their cognitive resources to the studied topic. At the same time, while the CM creator focuses on the concept mapping technique²². This technique avoids the cognitive overload that could be generated by requiring the student to learn to produce the CM while learning a new content to be mapped, facilitating the teaching-learning process²³. The CMs produced in our study were reviewed to ensure there was no saturation of information and the inclusion of unnecessary or confusing visual information, avoiding cognitive overload²³.

After the Concept Maps (CMs) production, a propositional clarity table was used to correct the CMs semantically. This step allowed for the analysis of each proposition individually to verify its clarity and precision². Thus, conceptual errors were corrected. Through the establishment of the Focus Question (FQ), hierarchical organization of concepts, semantic clarity of propositions, and continuous revisions, the necessary parameters for the proper production of CMs were followed²¹. Consequently, the main quality indicators of a CM were considered, such as its structure and content, which refer to the graphical organization and the set of propositions, respectively²³.

The CMs developed in this study were constructed through systematic searches in the scientific literature, which increases their reliability. However, it is important to recognize that CMs can be improved and refined over time as new scientific evidence emerges.

An integrative literature review gathered studies that showed the advantages of using CMs as a learning resource that allows for the organization of knowledge, the hierarchy of concepts, the development of students' autonomy, and critical thinking²⁰. Studies showed more meaningful learning, with more significant cognitive gain and integration between theory and practice, through CMs compared to traditional teaching methods, such as discussion based on questions and answers²⁴, for students in health-related courses^{25,26}. However, learning the concept mapping technique and putting it into practice can initially seem difficult²⁷, in addition to being a task that requires practice and time²⁸. Thus, students may initially have difficulties adhering to concept maps (CMs). However, with the inclusion of a facilitator whose role is to discuss the peculiarities of CMs and explain how to use the tool, these difficulties are gradually overcome²⁷. Therefore, it is suggested that CMs be integrated into the educational process, as they are highly effective in improving students'

academic performance²⁸. With this strategy in mind, our study opted to develop CMs to provide them ready-made to students.

As limitations of this study, it was observed that the number of studies on CMs in Dentistry, especially in clinical pharmacology, is still tiny. Few have compared the effectiveness of using CMs with traditional pedagogical techniques in Dentistry students. Furthermore, there are not many clinical studies, both observational and experimental, on the impact of using antithrombotics in Dentistry, whose methodological quality is not always adequate. This fact explains the low number of systematic reviews produced on the subject, the quality of the evidence generated, and the low degree of recommendation of practices. Unlike the medical field, expert opinions are used to guide the management of patients using antithrombotics in dentistry. However, it is important to highlight that there are several particularities in dental procedures and, in most cases, medical studies cannot be used as a parameter for dental practice. Finally, although it was not the scope of this study, the need for the development of clinical research on the impact of the use of antithrombotics in Dentistry was observed, considering the bleeding risks associated with various types of procedures, hemostasis management, and the risk of drug interactions.

The study in question is preliminary and focused on constructing CMs based on scientific evidence. Future studies will be conducted to test the effectiveness of the developed CMs and compare them with other pedagogical strategies involving the addressed topic.

CONCLUSION

The produced CMs contain essential information for the training of Dentistry students on a topic of great importance, especially for surgical cases. Future studies will be conducted to test the effectiveness of this teaching strategy.

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