Iatrogenic wounds: a common but often overlooked problem

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Historical evolution of iatrogenic injury and iatrogenic wounds

The historical evolution of iatrogenic wounds is shown in Fig. 1 [5, 10–15]. The term iatrogenesis means “brought forth by a healer” and is derived from the Greek ἅτρός (iatros, “healer”) and γένεσις (genesis, “origin”), so it could refer to good or bad effects. Since at least the time of Hippocrates, people have recognized that a healer could cure diseases but also cause potential damage [10]. X-rays were discovered by Roentgen in 1895, and radiation-induced skin damage was reported the following year [11]. Stacher first reported skin necrosis due to anticoagulant therapy [12, 16]. Warfarin-induced skin necrosis almost always occurs by day 10 after therapy [17]. Since the 1950s, with the use of new biological materials, exposure to implants has increased [13]. Since 1953, the terms “iatrogenic” and “trauma” have appeared in an increasing number of reports [14]. In recent years, the incidence of iatrogenic wounds has increased [1]. The reasons for this increase are as follows [15, 18–21]: (1) human life has gradually prolonged, due to which the population of elderly people is increasing; (2) metabolic diseases are becoming increasingly more common; (3) surgical indications are expanding, more difficult operations are being performed, and operation time is getting longer; (4) new drugs (e.g., antitumor treatments, immune treatments, and hormones) and various types of implantable biological materials are being used; (5) drug-resistant bacteria have emerged; and (6) new therapeutic modalities, such as those involving electricity, magnetism, and light, are being developed.

Characteristics of iatrogenic wounds

Iatrogenic wounds come under the purview of all hospital departments. Subcutaneous and deep artificial implants can result in wounds, so the incidence of iatrogenic wounds is increasing in surgical departments, especially in the fields of cardiac surgery, neurosurgery, orthopedics, and plastic surgery, which have witnessed the extensive use of biological materials, an expansion of surgical indications, and continual increases in the age limit for surgery [22, 23]. Surgeries for infants and young children are also becoming more common. Despite the advancements in medical science, the incidence of sternum infection and dehiscence after thoracotomy has remained unchanged. Skin and soft tissue necroses usually occur due to improper dressing, radiation therapy, or infusion treatment in oncology. The “cupping” therapy of traditional Chinese medicine and external treatment with herbs can lead to accidental burns and refractory wounds [24].

The consequences of some iatrogenic wounds are serious. If these wounds are not treated correctly, they may cause serious consequences and even death. For example, wounds caused by surgical implants may not show signs of infection even though many bacteria breed around surgical implants and present a clone-like growth pattern, and can even cause death.

Classification of iatrogenic wounds

Like all wounds, iatrogenic wounds can be classified into acute, chronic, and refractory wounds, depending on the duration of the wound. According to the level of difficulty of treatment, iatrogenic wounds can be divided into simple wounds and complex or refractory wounds [25–27]. Additionally, iatrogenic wounds can also be divided into clean wounds, clean-contaminated wounds, contaminated wounds, and infected wounds.

The causes of some iatrogenic wounds are clearly defined and can be easily identified, such as complications of surgery. However, there exist also some less obvious iatrogenic wounds, such as wounds caused by complex...
drug interactions, which may be identified through careful and detailed research.

Iatrogenic wounds can also be divided into avoidable and unavoidable wounds. Unavoidable iatrogenic wounds are necessarily caused by the treatment itself, such as secondary wounds of postoperative laser stripping treatment for pigment disease and donor site wounds after skin and flap grafting in plastic surgery. Clean surgical wounds tend to heal without complications. Avoidable iatrogenic wounds include various interventions in medical practice, like implants or materials, side effects of drugs, and medical errors.

Iatrogenic wounds are not caused only by surgeons but can be caused by almost any healthcare professional, including physical therapists, radiation technicians, dermatologists, community doctors, laser therapists, and nurses. Furthermore, iatrogenic wounds are not associated with only modern medicine (e.g., implants, radiation meters, and electric knives), but can result from traditional medicine as well (e.g., topical traditional Chinese medicines, cupping, and moxibustion). Iatrogenic wounds can be caused by the increased use of new tissue substitutes, new photoelectric instruments, and new chemotherapy drugs as well as the expanding indications for treatments (e.g., increased patient-age range and basic diseases such as diabetes/high blood pressure control).

The classification of iatrogenic wounds is different from that of other wounds dependent on the cause of the pathogenic factors. According to the pathogenic factors involved, iatrogenic wounds may be divided into wounds caused by SSIs; wounds caused by radioactive damage; wounds caused by lasers, electric coagulation, or electric knives; and wounds caused by drugs. SSIs represent the second most common cause of hospital-acquired infections and the most common type of healthcare-associated infection and substantially contribute to annual morbidity, healthcare costs, and mortality [28–31]. Iatrogenic wounds caused by SSIs are often difficult to treat.

In recent years, wounds induced by implanted materials are the most common type of iatrogenic wounds, and these tend to be intractable (Fig. 2) [28–34]. It is estimated that the annual rate of infections associated with surgical implants could be close to one million [35]. Overall, 2.6 million patients receive orthopedic prostheses in the USA each year, and the number of infections related to orthopedic prostheses is close to 112,000 (about 4.3%) [36]. The infection rate after joint replacement is 1% to 10%, depending on the surgery type and technique employed, body location, and aftercare [37–39]. Vascular surgery and groin surgery are associated with a high rate of SSIs [40]. Moreover, biofilm-related infections caused by *Staphylococcus aureus* are increasingly being detected in patients receiving intravascular catheters, cardiac pacemakers, vascular grafts, mechanical heart valves, and orthopedic implants [41, 42].

The number of patients with cancer has been increasing sharply worldwide each year [43]. Many cancer patients need radiotherapy, and the radiation time and dose are also increasing. Although accurately estimating radioactive damage rates is difficult, the incidence of iatrogenic radioactive skin wounds has increased significantly.

**Prevention of iatrogenic wounds**

Attention must be paid to pre-existing diseases, surgical time, wound contamination, patient age, malignant tumors, metabolic disease, malnutrition, immune suppression, smoking, etc. [44]. During surgery, frequent changes in the patient's position must be reduced. Reasonable application of surgical instruments must be ensured, and we should increase antimicrobial treatment appropriately for patients with longer operation times or excessive blood loss. Changes in the patient's position should be gentle during surgery to avoid damage to tissue. Radical removal of necrotic tissues in surgical sites must be ensured to prevent the formation of dead space. Close monitoring of body temperature is necessary during surgery to avoid temperature anomalies. Adequate intake of nutrients for patients must be ensured. The technique of fractionation of doses is used to minimize the risk of injury to normal tissue during radiation treatments [45]. Wound healing monitoring is an important concern in all surgical procedures since it allows the identification of signs or/and symptoms possibly related to surgical complications [46].

**Treatment of iatrogenic wounds**

The principles of treatment of iatrogenic wounds are the same as those for other wounds, though the former do have their own unique features. However, because iatrogenic wounds are caused by medical activities, patients are often reluctant to cooperate or psychologically fear and are reluctant to accept more traumatic treatments. Medical staff must pay attention to the psychological treatment of patients to avoid complaints and emotional disturbance. Thus, the treatment of iatrogenic wounds while avoiding secondary injuries is a challenge for medical staff.

The pathway of healing is determined by characteristics of the wound on initial presentation, and it is vital to select the appropriate method to treat the wound based on its ability to avoid hypoxia, infection, excessive edema, and foreign bodies [47]. It is relatively simple to treat wounds that are unavoidable, such as wounds after laser treatment and donor site wounds after skin graft removal. These wounds should be kept clean and dry, and steps must be taken to reduce exudation and prevent infection; with these measures, most of these wounds heal without complications. For simple wounds, infection should be controlled to prevent wound deepening; most of these wounds heal in 1 to 2 weeks. For
complex or refractory wounds, it is necessary to choose a comprehensive treatment based on the condition of the wound. These wounds may require various treatment strategies, including nutritional support, exogenous growth factors, chitosan, hyperbaric oxygen, platelet concentrate, exogenous alginate or biological dressings, debridement, and surgery [48, 49].

Since wounds caused by SSIs account for a large proportion of iatrogenic wounds, guidelines have been developed for the prevention and treatment of SSIs. The treatment of SSIs includes a variety of comprehensive treatments such as pre-hospital interventions, hospital interventions, and post-discharge incision care [50].

In the case of non-iatrogenic wounds, any foreign bodies present within the wound must be removed. Similarly, in the case of implant-related iatrogenic wounds, the implant should be removed and then replaced 4 to 6 months later. However, in some cases, the implant is expensive or essential to the patient, such as pacemakers, silicone breast implants, artificial vascular grafts, and periprosthetic joints [51, 52]. In such cases, salvage treatment can be performed, and if necessary, a salvage operation should be performed to preserve the implants as much as possible and minimize the damage to the patient. Debridement and prosthesis retention may bring good quality of life outcomes to patients and reduce costs [53]. Byren et al. [54] showed that the success rate of 112 infected arthroplasties treated with debridement, antibiotics, and implant retention was 81%. A systematic literature review by Maillet et al. reported that debridement and prosthesis retention in association with prolonged antimicrobial treatment may be an advantageous alternative to arthroplasty exchange for frail patients [55].

A review of the literature showed that the treatment of implant-related iatrogenic wounds usually includes the following [44, 52, 56–58] (Fig. 3): (1) the control of systemic infection; (2) local debridement to remove necrotic tissue; (3) wound cleaning and debridement to
retain implants, followed by repeated rinsing with a high-pressure washing gun, hydrogen peroxide, and saline, and finally negative-pressure wound therapy; and (4) a well-vascularized myocutaneous flap to cover the wound. When no suitable tissue is present around the wound, the prosthesis can be enclosed with a capsule. Secondary closure of these wounds is usually successful in patients with no related systemic diseases, and sufficient and well-vascularized soft tissue coverage. Successful salvage of ophthalmic and breast implants in patients with infected wounds has been achieved using the above method in China and other countries [57, 58]. Postoperative observation is necessary for the prevention and control of hematoma, infection, and skin flap necrosis. Additionally, proper management is indispensable for tetanus-prone wounds. However, avoiding the recurrence of implant infection is difficult, and implant removal is inevitable in some cases. Vacuum sealing drainage may be applied to enable subsequent wound coverage with a skin graft or skin flap.

Conclusion
Iatrogenic wounds are a common problem with unique features. Medical staff must be better educated on medical ethics and improve their medical knowledge to avoid the occurrence of avoidable iatrogenic wounds. If iatrogenic wounds do occur, efforts must be made to accelerate wound healing as soon as possible while avoiding secondary injuries.

Abbreviations
SSI: Surgical site infection

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Authors’ contributions
BC reviewed the literature, interpreted the information, and drafted the review. JT and YP helped create the figures, compiled the reference list, and revised the manuscript as well as subsequently updated it as appropriate. FX provided the recommendations and helped draft and revise the review. All authors read and approved the final manuscript.

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Not required

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Competing interests
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