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Advancements and trends in digestive system autotransplantation: a bibliometric and visualization analysis

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Background: Digestive system autotransplantation is an emerging surgical technique used to treat complex digestive diseases.

Methods: This study conducted a bibliometric analysis of 748 publications from the Web of Science Core Collection (WoSCC) database, using VOSviewer and CiteSpace tools to map research trends, author contributions, and institutional collaborations. Articles were selected based on their relevance to digestive system autotransplantation, focusing on autologous liver, pancreatic, and small intestine transplantation. The analysis included publication volume, citation counts, key authors, leading journals, and keyword co-occurrence.

Results: The analysis revealed a steady rise in publications between 2004 and 2015, followed by a gradual decline after 2016. The United States leads in research output, accounting for 40.11% of publications, followed by China. The leading institutions are predominantly based in the United States, with the University of Minnesota System producing the most publications. High-frequency keywords include autologous transplantation, pancreatic transplantation, chronic pancreatitis (CP), and postoperative complications.

Conclusion: Digestive system autotransplantation is a promising approach for complex cases. Continued interdisciplinary collaboration and focus on clinical outcomes will drive future advancements.

KEYWORDS

digestive system autotransplantation, autologous transplantation, pancreatic transplantation, chronic pancreatitis, bibliometric analysis

Introduction

Digestive system autotransplantation has emerged as an advanced surgical technique, demonstrating significant potential in treating complex digestive system diseases in recent years. This technique includes autologous liver transplantation, pancreatic transplantation, and small intestine transplantation (1-3). Traditional digestive surgeries, such as liver resection, pancreatic resection, and small intestine resection, have achieved some success in treating tumors and other diseases. However, for tumors located in complex anatomical

areas adjacent to critical blood vessels, the surgical risks and difficulties are extremely high, often rendering them "unresectable" (4, 5). These unresectable tumors, due to their special locations, make it challenging to completely remove the tumor without damaging vital structures during surgery, thus increasing the risk of postoperative recurrence and complications (6). Advances in surgical techniques, such as vascular occlusion, vena cava shunting, and *ex vivo* resection, have made autotransplantation feasible (7). This technique combines traditional resection and organ transplantation, involving the removal, repair, and reconstruction of the entire organ *ex vivo* in cold preservation, and then reimplanting the unaffected part of the organ (8–10). The development of this technique not only provides new treatment options but also offers surgeons more choices when dealing with complex cases.

Autologous liver transplantation combines the advantages of traditional liver resection and liver transplantation. It provides a new treatment strategy, especially for complex conditions such as multiple liver tumors or hepatic echinococcosis (2, 11). Compared to whole liver transplantation, autotransplantation does not require a donor liver, avoiding the risks of donor shortage and long-term postoperative immunosuppression, while also reducing the risk of disease progression and waiting time for a donor (12). This technique allows for more precise removal of lesions, reducing the risk of uncontrollable bleeding during surgery, increasing the surgical space for complex vascular reconstruction, and minimizing ischemic damage. Despite these advantages, digestive system autotransplantation is highly challenging, requiring a highly skilled surgical team, and has not yet been widely standardized in clinical practice (13, 14). This partly explains why this technique has not been widely adopted. Successful implementation of autotransplantation requires a surgical team with high professional skills and extensive experience, along with precise preoperative assessments and intraoperative support. However, with the development of preoperative three-dimensional image reconstruction technology and liver indocyanine green clearance technology, preoperative estimation of the remnant liver volume has become possible (15-17). This pre-assessment technology, through precise measurement and simulation of intraoperative conditions, greatly improves the safety and success rate of the surgery (11, 18, 19). Hepatic alveolar echinococcosis (HAE) is a severe parasitic infection often leading to extensive liver tissue destruction. Traditional surgery struggles to completely remove the diseased tissue, while autotransplantation provides a new possibility for radical treatment (11). Furthermore, in treating hepatobiliary malignancies, autotransplantation also shows significant advantages by completely removing lesions and preserving as much healthy tissue as possible, significantly improving patient prognosis (20).

Autologous pancreatic transplantation shows unique advantages in treating chronic pancreatitis (CP) and preventing postoperative diabetes (21). Chronic pancreatitis and pancreatic tumors pose significant therapeutic challenges due to their complex anatomical structures and the essential endocrine function of the pancreas (22). Traditional surgical interventions, such as pancreaticoduodenectomy or distal pancreatectomy, can relieve symptoms and remove tumors but often result in substantial loss of pancreatic function, leading to diabetes (23). Autologous pancreatic transplantation, particularly autologous islet transplantation, offers a promising alternative. This technique involves isolating islet cells from the resected pancreas and then autotransplanting them into the liver or other suitable locations in the patient's body. This method helps preserve endocrine function and reduce the risk of postoperative diabetes (21). Autologous islet transplantation is particularly beneficial for patients undergoing total pancreatectomy (TP) for chronic pancreatitis, as it can significantly improve their quality of life by maintaining insulin independence or reducing insulin demand (24, 25). The technical complexity of autologous pancreatic transplantation lies in the isolation and transplantation of islet cells. Advances in islet isolation techniques and immunosuppressive regimens have improved the outcomes of this surgery, making it a viable option for a broader range of patients (25). Preoperative planning, including imaging studies and functional assessments, is crucial for optimizing results and reducing complications.

Autologous small intestine transplantation is primarily used to treat extensive small intestine diseases, such as short bowel syndrome (SBS) and certain complex intestinal tumors (26, 27). The small intestine is crucial for nutrient absorption, and its extensive resection can lead to short bowel syndrome (SBS), characterized by malnutrition, diarrhea, and severe nutritional deficiencies (28). Traditional surgical treatments for SBS are usually limited and may not restore sufficient intestinal function (29). Autologous small intestine transplantation has emerged as a new approach to address these challenges. This surgery involves the resection of the affected small intestine, ex vivo treatment, and reimplantation into the patient's body. By preserving and reusing the patient's own small intestine, this technique aims to restore the continuity and function of the intestine, thereby improving the patient's nutritional status and quality of life. Advances in surgical techniques and perioperative care have enhanced the feasibility and effectiveness of autologous small intestine transplantation. Preoperative evaluation, including detailed imaging studies and functional tests, is crucial for ensuring surgical success. Additionally, postoperative care, including nutritional support and monitoring of complications, plays a key role in the long-term success of this approach (30, 31).

In summary, digestive system autotransplantation, as a cuttingedge surgical technique, is gradually becoming an important means of addressing complex digestive system diseases. Through continuous technological innovation and accumulation of clinical practice, this technique is expected to be more widely applied in the future, bringing hope for recovery to more patients. With the strengthening of global medical exchanges and cooperation, the promotion and standardization of autotransplantation technology will be further enhanced, ultimately benefiting a broad range of patients. Although research on digestive system autotransplantation is relatively in-depth, quantitative data is still lacking. Despite some scholars discussing studies with conflicting data through meta-analyses, the depth of discussion is relatively limited and subjective (32-34). Moreover, many scholars' research on digestive system autotransplantation remains at the level of literature review and personal clinical experience summaries (9, 35).

Bibliometrics is a tool for quantitatively analyzing literature, which can reveal research hotspots, frontier dynamics, and research trends in a particular field (36). Through statistical analysis of relevant literature, we can understand the research activity, main contributors, and collaboration relationships among research institutions in the field, thereby providing a scientific basis for future research. Bibliometrics, through quantitative analysis of a large amount of literature data, helps researchers identify and evaluate scientific research trends and patterns. It typically involves several analyses: counting publications over time to gauge research activity and trends; identifying research hotspots through keyword and subject term co-occurrence; evaluating impact via citation analysis; and examining collaboration networks to understand relationships among research institutions and authors. Through these analyses, we can systematically review the research status of the field of digestive system autotransplantation, identify key researchers and institutions, understand research hotspots and trends. This comprehensive overview will provide valuable insights for future research directions and promote better planning and collaboration within the scientific community.

In this study, we aim to systematically review and identify the key areas of focus and emerging patterns in the field of digestive system autotransplantation. Using visualization tools VOSviewer and CiteSpace, we reviewed and analyzed relevant literature from 2004 to 2024. We conducted a systematic analysis of the current state and evolving patterns in digestive system autotransplantation using bibliometric methods. Through these analyses, we can systematically review the research status of the field of digestive system autotransplantation, identify key researchers and institutions, understand research hotspots and trends. This comprehensive analysis will offer important perspectives for guiding future research directions and promote better planning and collaboration within the scientific community.

Methods and materials

The Web of Science Core Collection (WoSCC) database is known for its precise document type labeling, making it ideal for literature analysis. Thus, we selected this database for our search (37, 38). On May 22, 2024, we conducted a search in WoS for articles on digestive system autotransplantation published between 2004 and 2024. To ensure comprehensive coverage and minimize the risk of missing relevant literature, the search strategy incorporated a wide range of synonyms and related terms for "autotransplantation" and the specific digestive organs. The detailed search strategy, including the combination of keywords and operators used, is presented in Supplementary Table S1. The search formula included: (((((((TS=(Autotransplantation)) OR TS=("Autologous Transplantation")) OR TS=("Autologous Transplantations")) OR TS=("Transplantations, Autologous")) OR TS=(Autografting)) OR TS=(Autograftings)) OR TS=(Autotransplantation)) OR TS=(Autotransplantations) AND (((((((((TS=(Liver)) OR TS=(Livers)) OR TS=(Pancreas)) OR TS=(pancreatic)) OR TS=("Intestine, Small")) OR TS=("Intestines, Small")) OR TS=("Small Intestines")) OR TS=("Small Intestine"))) OR TS=(intestine)) OR ((((((((TS=("autologous liver



transplant")) OR TS=("autologous liver transplantation")) OR TS=("Autologous Pancreas Transplantation")) OR TS=("Autologous Pancreas Transplant")) OR TS=("autologous small intestine transplant")) OR TS=("Autologous transplantation")) OR small bowel TS=("intestinal autotransplantation")) OR TS=("Pancreas autotransplantation")) OR TS=("intestinal autotransplant")) OR TS=("Pancreas autotransplant")) OR TS=("Liver autotransplantation")) OR TS=("liver autotransplant").

The criteria for selecting literature for this study were: (1) full-text publications on digestive system autotransplantation; (2) articles and reviews in English; and (3) publications from 2004 to 2024. Exclusion criteria included: (1) unrelated topics; and (2) conference abstracts, news, brief reports, etc. Plain text versions of the papers were extracted.

We utilized GraphPad Prism v8.0.2 for analyzing and plotting annual and national publication trends and proportions. Additionally, CiteSpace (v6.2.4R) and VOSviewer (v1.6.18) were employed to analyze the data and create visual representations of the scientific knowledge maps. Figure 1 shows the flow chart of the literature selection process.

Results

From 2004 to 2024, the WoSCC database recorded a total of 748 publications on autologous transplantation in the digestive system, comprising 618 research articles and 130 reviews. These publications involved 49 different countries, 943 institutions, and 3,822 authors.



Since 2004, the annual number of published papers has shown a gradual rise (see Figure 2). This trend can be segmented into three phases: between 2004 and 2007, the yearly publication count was around 20, indicating slow growth and limited research interest; from 2008 to 2015, there was a steady increase in publications, peaking in 2015; post-2016, the number of publications started to decline gradually.

Countries and institutions

Research on autologous transplantation in the digestive system spans 49 countries. Figures 3A, B illustrate the annual publication volume for the top 10 countries over the past decade. The leading five countries in this field are the United States, China, Italy, Japan, and the United Kingdom. The United States is the frontrunner, accounting for 40.11% of the total publications, significantly outpacing other countries.

Among the top 10 countries for paper publications, the United States stands out with a citation count of 7,323 (see Supplementary Table S2), far surpassing other nations. Additionally, its citation-to-publication ratio is 24.41, ranking sixth overall, which indicates the high quality of its publications. China is second with 172 publications and 2,561 citations, but its citation-to-publication ratio is relatively low at 14.89. Figure 3C depicts the collaboration network, showing close cooperation between the United States and China. The U.S. also collaborates extensively with Italy, the United Kingdom, and France, while China has strong partnerships with Japan, Germany, and South Korea. The United States not only excels in publication volume and citation frequency but also has a centrality score of 0.54, underscoring its leading role in the field.

In total, 943 institutions have published articles on gastrointestinal autotransplantation. Of the top ten institutions by publication volume, eight are based in the United States, and two are located in China (refer to Supplementary Table S3 and Figure 4). The University of Minnesota System has the highest number of publications with 71 papers, cited 2,258 times, averaging 31.80 citations per paper. The University of Minnesota Twin Cities ranks second with 70 papers, cited 2,101 times, averaging 30.01 citations per paper. The University System of Ohio ranks third with 31 papers, cited 350 times, averaging 11.29 citations per paper. Cincinnati Children's Hospital Medical Center is ranked fourth, with 23 papers cited 227 times, averaging 9.87 citations per paper. Further analysis shows that institutions, both domestic and international, tend to collaborate more with others within their own countries. Therefore, we encourage increased collaboration between domestic and international institutions to break down academic barriers.

Journal analysis and co-citation data

Supplementary Tables S4, S5 detail the top 10 journals by publication volume and citation counts. The American Journal of Transplantation (30 papers, 4.01%) is the leading journal in this field, followed by Transplantation Proceedings (30 papers, 4.01%), and Transplantation (19 papers, 2.54%; Figure 5A). Of the top 10 prolific journals, Annals of Surgery boasts the highest impact factor (IF) of 10.1. Notably, 70% of these journals fall within the Q1 or Q2 categories.

A journal's influence is measured by the frequency of its cocitations, reflecting its importance within the scientific community. As shown in Figure 5B and Supplementary Table S5, the most frequently co-cited journal is "Annals of Surgery" (362 times), followed by "Transplantation" (338 times) and "Surgery" (300 times). Among the 10 most frequently co-cited journals, the "New England Journal of Medicine" stands out with 258 citations and the highest impact factor of 158.5. Notably, 80% of the co-cited journals fall within the Q1/Q2 quartiles.

The thematic distribution of academic publications is depicted using dual-map overlays (Figure 5C). Colored paths illustrate citation relationships, with citing journals on the left and cited journals on the right. The results reveal two primary citation paths: research published in journals related to molecular/biology/genetics is mainly cited by journals in molecular/biology/immunology and medicine/medical/clinical fields. Additionally, research published in journals related to health/nursing/medicine is primarily cited by journals in the medicine/medical/clinical fields.

Among all authors who have published literature on digestive system autotransplantation, Supplementary Table S6 lists the top 10 authors with the highest number of publications. These leading authors have collectively contributed 223 papers, accounting for 29.81% of all publications in this field. Bellin, Melena D., stands out with the most research papers at 43, followed by Freeman, Martin L., with 23 papers. Chinnakotla, Srinath, Naziruddin, and Pruett, Timothy L. each have published 22 papers. CiteSpace provides a visualization of the collaborative network among these authors (Figure 6A).

Figure 6B and Supplementary Table S6 show the top 10 most co-cited and cited authors. In total, 146 authors have been cited more than 50 times, indicating the substantial influence and





reputation of their research. The most prominent nodes in the cocitation network are Bellin, Melena D. (171 citations), Sutherland, D.E.R. (163 citations), and Chinnakotla, Srinath (100 citations). Further analysis reveals that Bellin, Melena D. ranks first in both the number of publications and citations, underscoring his significant impact and leadership in the field of digestive system autotransplantation.

These findings highlight the contributions of key researchers and their collaborative networks, providing insights into the influential figures and their connections within the field. The data suggests that a small group of prolific authors and highly-cited researchers drives much of the advancement in digestive system autotransplantation, indicating potential focal points for future research collaborations and studies.

Co-cited references

The co-cited reference network, constructed using 1-year intervals from 2004 to 2024, consists of 805 nodes and 3,152 links (Figure 7A). Based on the top 10 most co-cited articles (Supplementary Table S7), the study "Total Pancreatectomy and Islet Autotransplantation for Chronic Pancreatitis," published in the Journal of the American College of Surgeons, reports that total pancreatectomy (TP) combined with intraportal islet autotransplantation (IAT) can alleviate pain and preserve a significant number of β -cells in patients with chronic pancreatitis (CP) who are unresponsive to other treatments.

From February 1977 to September 2011, 49 CP patients, including 53 children aged 5–18 years, underwent TP-IAT. The causes of CP were as follows: idiopathic (41%), sphincter of Oddi dysfunction/biliary (9%), genetic factors (14%), pancreas divisum (17%), alcohol (7%), and other causes (12%). The average age was 35.3 years, with 74% of the patients being female. Prior surgeries included Puestow (9%), Whipple (6%), distal pancreatectomy (7%), and other surgeries (2%). Post-surgery, islet function was categorized into three groups: insulin-independent for those not using insulin, partial islet function for those with hyperglycemia managed by daily insulin or C-peptide positive, and insulin-dependent for those requiring standard insulin therapy.

Starting in 2007, patients completed the SF-36 quality of life survey before treatment and during continuous follow-ups, with an additional comprehensive survey added in 2008. The actuarial survival rates post TP-IAT were 96% for adults and 98% for children at 1 year, and 89% and 98% at 5 years, respectively. Complications requiring reoperation occurred in 15.9% of patients, with bleeding being the most common at 9.5%. IAT function was achieved in 90% of patients (C-peptide >0.6 ng/ml). After 3 years, 30% of patients were insulin-independent (25% of adults and 55% of children), and 33% had partial islet function. Additionally, 82% had an average HbA1c below 7.0%. Prior pancreatic surgeries were associated with reduced islet yield (2,712 vs. 4,077/kg; p = 0.003). Islet yield categories correlated with function: <2,500/kg (36%),







2,501-5,000/kg (39%), >5,000/kg (24%), with corresponding 3year insulin dependence rates of 12%, 22%, and 72%, and partial function rates of 33%, 62%, and 24%.

All patients reported experiencing pain before TP-IAT, with nearly all using narcotics daily. Post-treatment, 85% reported improved pain levels, and after 2 years, 59% had ceased using narcotics. Among children, 94% reported pain improvement, with 67% being pain-free, although 39% continued using narcotics at follow-up. SF-36 survey results showed significant improvement in all dimensions, both physical and mental, compared to baseline (p < 0.01). TP can effectively relieve pain and improve the quality of life for refractory CP patients, even if complete narcotic withdrawal is delayed or incomplete. IAT preserves islet function in most patients, with over two-thirds maintaining function and a quarter of adults and half of children achieving insulin independence.

The second most cited article, "Total Pancreatectomy and Islet Autotransplantation in Children for Chronic Pancreatitis: indications, surgical techniques, postoperative management, and long-term outcomes," published in Annals of Surgery by Chinnakotla, Srinath, details the surgical techniques, complications, and long-term outcomes of TP-IAT. Surgical treatment for pediatric pancreatitis remains challenging; while partial resection or drainage surgery often provides temporary pain relief, long-term recurrence is common due to diffuse pancreatic involvement. TP eliminates the pain source, and IAT potentially prevents or mitigates TP-related diabetes. A retrospective study conducted from 1989 to 2012 reviewed 75 children with CP who underwent TP-IAT after failing medical, endoscopic, or previous surgical treatments. Following TP-IAT, 90% of patients showed significant improvement in pancreatitis pain and severity (p < 0.001), with sustained narcotic relief. Of the 75 patients, 31 (41.3%) achieved insulin independence. Factors associated with insulin independence included younger age (p = 0.032), absence of prior Puestow procedure (p = 0.018), smaller body surface area (p = 0.048), higher islet equivalents per kilogram (IEQ; p = 0.001), and a total IEQ > 100,000 (p = 0.004).

Multivariable analysis identified three significant factors associated with post-TP-IAT insulin independence: male gender, smaller body surface area, and higher total IEQ per kilogram. The total IEQ >100,000 was the single most significant factor associated with insulin independence (odds ratio = 2.62; p < 0.001). β -cell function was dependent on islet yield. The study concludes that TP-IAT is an effective treatment for children with painful pancreatitis unresponsive to medical and/or endoscopic treatments, providing significant long-term pain relief and improving quality of life.

We performed co-cited reference clustering and temporal clustering analysis, as shown in Figures 7B, C. Our findings reveal that intramuscular (cluster 4), embryonic (cluster 6), islets (cluster 8), and pig (cluster 9) were early research hotspots. Mid-period research hotspots included islet isolation (cluster 3), bile duct (cluster 11), cardiomyogenic differentiation (cluster 12), dexmedetomidine (cluster 13), pancreaticojejunostomy (cluster 14), cohort (cluster 15), MR imaging (cluster 16), mesenchymal

stromal cell (cluster 17), and pancreatoduodenectomy (cluster 18). Current and trending topics in the field are pancreatectomy (cluster 0), islet transplantation (cluster 1), hepatic alveolar echinococcosis (cluster 2), diagnosis (cluster 5), type 3c diabetes mellitus (cluster 7), and beta cells (cluster 19).

Using VOSviewer to examine keyword co-occurrence, we identified the most frequently used keywords: "complications" (163 occurrences), among others (Supplementary Table S8, Figures 8A, B). We excluded unrelated keywords and built a network of 172 keywords that appeared at least seven times, resulting in five distinct clusters. Cluster 1 (pink) contains 76 keywords including autologous transplantation, angiogenesis, stem cell, diabetes, cell transplantation, survival, preservation, blood, T cell, regeneration, apoptosis, oxidative stress, reperfusion injury, inflammation, beta cells, bone marrow, cold storage, differentiation, donor, failure, gene therapy, ischemia reperfusion injury, pig, model, pancreatic islets, and pathogenesis. Cluster 2 (green) includes 37 keywords such as total pancreatectomy, pain, chronic pancreatitis, quality of life, islet autotransplantation, long-term outcome, exocrine insufficiency, efficacy, follow-up, management, multicenter, and head resection. Cluster 3 (purple) comprises 34 keywords including outcome, surgery, complication, resection, experience, hepatectomy, tumor, infection, graft, impact, mass, inferior vena cava, reconstruction, root, splenectomy, tissue, and implantation. Cluster 4 (yellow) contains 15 keywords such as acute autotransplantation, children, chronic pancreatitis, cystic fibrosis gene, independence, insulin, risk, mutation, total pancreatectomy with islet autotransplantation (TPIAT), yield, and hereditary pancreatitis. Cluster 5 (yellow) consists of 10 keywords including beta cell function, clinical research, glucose, insulin independence, islet isolation, islets of Langerhans, pediatrics, and practice. We used CiteSpace to create a volcano plot that visually displays how research hotspots have evolved over time (Figures 8C, D).

Co-cited references and keywords

Using CiteSpace, we identified the 50 most reliable citation bursts in the field of digestive system autotransplantation. All 50 references were published between 2004 and 2024, indicating their frequent citation over the past two decades. Importantly, 12 of these papers are currently at their peak citation period (Figure 9), suggesting that digestive system autotransplantation will continue to be a focal point in future research.

Among the 576 strongest burst keywords in this field, we focused on the top 50 with the highest burst strength (Figure 10). These keywords represent current research hotspots and potential future directions in the field.

Discussion

This study systematically retrieved and analyzed literature from the WoSCC database on autologous transplantation of the digestive system, comprehensively showcasing the research trends and academic influence in this field over the past two decades. Autologous transplantation of the digestive system is a significant surgical technique, particularly crucial in treating hepatic hydatid disease, chronic pancreatitis, and certain refractory digestive system diseases. This systematic review of the literature in this field helps understand its current research status, hotspots, and future development directions. The primary reason for selecting the WoSCC database for literature retrieval is its high accuracy in documenting literature types, providing comprehensive and detailed records. By retrieving relevant literature from 2004 to 2024, we ensured the completeness and timeliness of the data. The search strategy employed multiple keywords and synonyms to cover all relevant literature. The establishment of inclusion and exclusion criteria further enhanced the scientificity and rationality of literature screening, ensuring the quality and reliability of the data.

From the perspective of publication trends, since 2004, the number of publications on autologous transplantation of the digestive system has generally shown an upward trend, peaking between 2008 and 2015. This growth trend may reflect the continuous development of new technologies and treatment methods during this period. However, since 2016, the number of publications has declined, possibly due to a shift in research focus, the maturation of new technologies, and competition from other treatment methods. This change in trend suggests the need to pay attention to emerging hotspots and potential future development directions in this field.

Although the number of publications has decreased in recent years, it does not imply a reduction in the importance of autologous transplantation techniques for the digestive system. On the contrary, these techniques continue to play an indispensable role in clinical applications, providing solutions that other treatment methods cannot replace, especially in handling complex cases and improving patient survival quality. Therefore, future research should continue to focus on improving the operability and safety of these techniques, optimizing postoperative management strategies, and exploring new application scenarios to ensure patients receive the best treatment outcomes and quality of life.

Autologous liver transplantation is a crucial surgical technique, particularly suitable for patients who cannot undergo conventional liver transplantation. This technique is primarily used to treat diseases such as primary liver cancer, hepatic hydatid disease, and hepatic hemangioma (39, 40). A significant application of autologous liver transplantation involves liver resection, followed by *ex vivo* liver repair and subsequent reimplantation into the patient. Literature shows that autologous liver transplantation significantly improves patient survival rates and quality of life. However, the technique is complex, and the management of postoperative complications and long-term follow-up remain essential research directions.

Autologous pancreatic transplantation, mainly total pancreatectomy with islet autotransplantation (TP-IAT), plays a crucial role in treating chronic pancreatitis (41). This technique involves removing the diseased pancreas and transplanting healthy islet cells back into the patient, aiming to alleviate pain and preserve endocrine function. Studies indicate that TP-IAT not only significantly reduces patient pain but also enhances insulin independence and improves quality of life. However, improving the



Top 50 References with the Strongest Citation Bursts

References	Year	Strength Begin	End	2004 - 2024
Krause DS, 2001, CELL, V105, P369, DOI 10.1016/S0092-8674(01)00328-2, DOI	2001	4.53 2004	2006	
Ahmad SA, 2005, J AM COLL SURGEONS, V201, P680, DOI 10.1016/j.jamcollsurg.2005.06.268, DOI	2005	5.86 2007	2010	
Ryan EA, 2005, DIABETES, V54, P2060, DOI 10.2337/diabetes.54.7.2060, DOI	2005	5.07 2007	2008	
Clayton HA, 2003, TRANSPLANTATION, V76, P92, DOI 10.1097/01.TR0000054618.03927.70, DOI	2003	4.43 2007	2008	
Ahmad SA, 2006, CURR PROB SURG, V43, P127, DOI 10.1067/j.cpsurg.2005.12.004, DOI	2006	4.1 2007	2010	
Shapiro AMJ, 2006, NEW ENGL J MED, V355, P1318, DOI 10.1056/NEJMoa061267, DOI	2006	7.6 2008	2011	
Blondet JJ, 2007, SURG CLIN N AM, V87, P1477, DOI 10.1016/j.suc.2007.08.014, DOI	2007	11.5 2009	2012	
Sutherland DER, 2008, TRANSPLANTATION, V86, P1799, DOI 10.1097/TP.0b013e31819143ec, DOI	2008	9.92 2010	2013	
Webb MA, 2008, PANCREAS, V37, P282, DOI 10.1097/mpa.0b013e31816fd7b6, DOI	2008	7.7 2010	2013	
Argo JL, 2008, AM SURGEON, V74, P530	2008	6.04 2010	2013	
Rafael E, 2008, AM J TRANSPLANT, V8, P458, DOI 10.1111/j.1600-6143.2007.02060.x, <u>DOI</u>	2008	4.39 2010	2013	
Anazawa T, 2009, AM J TRANSPLANT, V9, P2383, DOI 10.1111/j.1600-6143.2009.02765.x, DOI	2009	5.29 2011	2014	
Garcea G, 2009, PANCREAS, V38, P1, DOI 10.1097/MPA.0b013e3181825c00, DOI	2009	4.76 2011	2014	
Sutton JM, 2010, SURGERY, V148, P676, DOI 10.1016/j.surg.2010.07.043, DOI	2010	6.68 2012	2015	
Sutherland DER, 2012, J AM COLL SURGEONS, V214, P409, DOI 10.1016/j.jamcollsurg.2011.12.040, DO	2012	21.16 2013	2017	
Bellin MD, 2011, CLIN GASTROENTEROL H, V9, P793, DOI 10.1016/j.cgh.2011.04.024, DOI	2011	8.36 2013	2016	
Bramis K, 2012, BRIT J SURG, V99, P761, DOI 10.1002/bjs.8713, DOI	2012	7.64 2013	2017	
Bellin MD, 2013, PANCREAS, V42, P317, DOI 10.1097/MPA.0b013e3182681182, <u>DOI</u>	2013	5 201 3	2016	
Ris F, 2011, TRANSPLANTATION, V91, P895, DOI 10.1097/TP.0b013e31820f0892, DOI	2011	5 2013	2016	
Morgan K, 2012, J GASTROINTEST SURG, V16, P129, DOI 10.1007/s11605-011-1744-y, DOI	2012	5 2013	2016	
Barton FB, 2012, DIABETES CARE, V35, P1436, DOI 10.2337/dc12-0063, <u>DOI</u>	2012	4.12 2013	2017	
Bellin MD, 2014, PANCREATOLOGY, V14, P27, DOI 10.1016/j.pan.2013.10.009, DOI	2014	7.26 2014	2019	
Lundberg R, 2013, AM J TRANSPLANT, V13, P2664, DOI 10.1111/ajt.12392, DOI	2013	4.66 2014	2016	
Bellin MD, 2012, CURR DIABETES REP, V12, P580, DOI 10.1007/S11892-012-0296-1, DOI	2012	4.4 2014	2017	
Chinnakotia S, 2014, ANN SURG, V260, P56, DOI 10.1097/SLA.000000000000569, DOI	2014	10.53 2015	2019	
Chinnakotia S, 2014, J AM COLL SURGEONS, V218, P530, DOI 10.1016/j.jamcoilsurg.2013.12.037, DOI 10.1016/j.jamcoilsurg.2014.12.037, DOI 10.1016.12.037, DOI 10.1016.12.037, DOI 10.1016.12.037, DOI 10.1016.12.037, DOI 10.1016.12.037, DOI 10.1016.12, DOI 1	2014	6.39 2015 E 4 201 E	2019	
Balzano G, 2013, ANN SURG, V258, P210, DOI 10.1097/SLA.00013631829C/900, DOI	2013	5.4 2015	2018	
Chinnakotia S, 2015, ANN SURG, V262, P610, DOI 10.1097/SLA.00000000000000000	2015	7 22 2016	2018	
Wilson GC, 2014, ANN SURG, V200, P039, DOI 10.1097/SEA.00000000000020, DOI	2014	1.52 2016	2010	
Pallia MD 2015, MANSPLANTATION, V95, P1051, DOI 10.1097/12.0001565162045100, DOI	2015	4.55 2016	2010	
Ali LIA 2012 APCH SURG, V201, P21, D01 10.1097/3LA.000000000000000000000000	2013	4.33 2016	2020	
Wilcov CM 2015 CLIN GASTROENTEROL H V13 P522 DOI 10.1016/j.coh.2014.10.015 DOI	2012	4.34 2010	2017	
Johnston PC 2015 L CLIN ENDOCR METAB V100 P1765 DOI 10.1210/jc.2014-4298 DOI	2015	4 23 2016	2020	
Wen H 2016 AM J TRANSPLANT V16 P615 DOI 10.1111/ait.13465 DOI	2016	11.09 2018	2021	
Bellin MD 2017 PEDIATR GASTR NUTR V64 P440 DOI 10 1097/MPG 00000000001314 DOI	2017	8 69 2018	2022	
He YB. 2015. J GASTROINTEST SURG. V19. P1457. DOI 10.1007/s11605-015-2842-z. DOI	2015	5.69 2018	2020	
Aii T. 2018. J HEPATOL, V69. P1037. DOI 10.1016/j.jhep.2018.07.006. DOI	2018	12.1 2019	2024	
Vang XW. 2018, AM J TRANSPLANT, V18, P1668, DOI 10.1111/ait.14621, DOI	2018	8.94 2019	2024	
Bellin MD, 2018. PANCREATOLOGY, V18. P286. DOI 10.1016/i.pan.2018.02.001. DOI	2018	7.76 2019	2024	
Lei JY, 2015, MEDICINE, V94, P0, DOI 10.1097/MD.000000000000514, DOI	2015	5.08 2019	2020	
Deplazes P, 2017, ADV PARASIT, V95, P315, DOI 10.1016/bs.apar.2016.11.001, DOI	2017	4.33 2019	2022	
Bellin MD, 2019, J AM COLL SURGEONS, V228, P329, DOI 10.1016/j.jamcollsurg.2018.12.019, DOI	2019	10.32 2020	2024	
Qiu YW, 2019, SURGERY, V165, P889, DOI 10.1016/j.surg.2018.11.007, DOI	2019	7.33 2020	2024	
Hwang R, 2018, J HEPATOL, V69, P1002, DOI 10.1016/j.jhep.2018.09.005, DOI	2018	5.49 2020	2024	
Zawistowski M, 2020, SURGERY, V168, P631, DOI 10.1016/j.surg.2020.05.036, DOI	2020	7.34 2021	2024	
Kato T, 2020, ANN SURG, V272, P766, DOI 10.1097/SLA.000000000004270, DOI	2020	4.66 2021	2024	
Kotagal M, 2019, AM J TRANSPLANT, V19, P1187, DOI 10.1111/ajt.15150, DOI	2019	4.14 2021	2024	
Abu-El-Haija M, 2020, PANCREATOLOGY, V20, P762, DOI 10.1016/j.pan.2020.04.005, DOI	2020	5.08 2022	2024	
Wen H, 2019, CLIN MICROBIOL REV, V32, P0, DOI 10.1128/CMR.00075-18, DOI	2019	4.76 2022	2024	
FIGURE 9 Top 50 references with the strongest citation bursts.				

survival rate and function of islet cells remains a primary research focus. Additionally, research has shown that the effectiveness of TP-IAT is closely related to factors such as the patient's age and the number of islets, necessitating further in-depth studies (42).

Autologous small intestine transplantation offers unique advantages in treating small intestine diseases and short bowel syndrome. This technique involves resecting the diseased small intestine and reimplanting healthy segments to restore normal small intestine function. Research indicates that autologous small intestine transplantation significantly improves nutrient absorption and reduces dependence on parenteral nutrition. However, the technique is challenging, and managing postoperative complications (such as graft dysfunction and infections) remains an important research direction. In recent years, advancements in surgical techniques and postoperative management have gradually increased the success rate of autologous small intestine

Top 50 Keywords with the Strongest Citation Bursts

Keywords	Year St	trength Begin	End	2004 - 2024
bone marrow	2004	5.64 2004 2	2012	
autologous transplantation	2004	4.77 2004 2	2014	
beta cells	2004	2.95 2004 2	2009	
bone marrow transplantation	2004	2.66 2004 2	2008	
nitric oxide	2005	3.65 2005 2	2008	
stem cells	2005	3.17 2005 2	2010	
preservation	2006	5.5 2006 2	2014	
angiogenesis	2008	2.82 2008 2	2012	
graft function	2009	4.46 2009 2	2013	
injury	2009	3.05 2009 2	2012	_
resection	2004	5.83 2010 2	2015	
auto transplantation	2005	5.1 2010 2	2013	_
in vitro	2004	5.08 2010 2	2014	
tissue	2010	3.63 2010 2	2015	
insulin	2008	2.56 2010 2	2014	
liver	2006	2.48 2010 2	2013	
donor	2011	3.05 2011 2	2012	_
cell transplantation	2008	2.82 2011 2	2016	
cells	2005	2.44 2011 2	2013	
pancreaticoduodenectomy	2010	3.08 2012 2	2017	
mice	2013	42013 2	2017	
inflammation	2013	2 92 2013 2	2016	
apontosis	2005	3 32 2014 2	2015	
nancreatogenic diabetes	2014	2 88 2014 2	015	
cancer	2014	2 57 2014 2	2016	
ischemia reperfusion iniury	2015	3 41 2015 2	2017	
expression	2015	2 03 2015 2	2016	
regeneration	2005	27 2015 2	2016	
type 1 diabates mellitus	2000	2.56 2015 2	2016	
long torm outers	2015	5.02 2015 2	010	
torm follow up	2015	2.59 2016 2	010	-
riele	2010	3.36 2016 2	000	-
risk intentional autotopografiantation	2010	2.80 2016 2	2020	
Intestinal autotransplantation	2010	2.0 2010 2	010	
isiets of langerhans	2016	2.55 2016 2	2017	_
risk factors	2012	4.38 2017 2	024	
experience	2011	5.49 2018 2	019	
nepatic alveolar echinococcosi	s 2018	5.21 2018 2	2024	_
children	2015	4.4 2018 2	2024	-
classification	2018	3.88 2018 2	2024	-
islet isolation	2011	3.49 2018 2	2021	_
clinical research	2016	3.02 2018 2	2021	_
islet transplantation	2008	2.99 2018 2	2021	
alveolar echinococcosis	2011	6.99 2019 2	2024	
inferior vena cava	2012	5.89 2019 2	2024	
reconstruction	2012	5.45 2019 2	2022	
diagnosis	2006	3.49 2019 2	2024	
outcm	2013	5.75 2020 2	2024	
type 1 diabetes	2012	3.1 2020 2	2021	
international study group	2020	2.76 2020 2	2024	
an when reception	2012	2 47 2020 2	2022	

Top 50 keywords with the strongest citation bursts.

FIGURE 10

transplantation, but further research is needed to optimize surgical protocols and enhance long-term survival rates.

From a national perspective, the United States has made the most significant contributions to research in this field, with the highest number of publications and citations, reflecting its leading position in autologous transplantation research for the digestive system. China, as the second-largest contributor, has seen a significant increase in research in recent years, demonstrating strong development momentum. Notably, a portion of China's research output addresses conditions like hepatic alveolar echinococcosis (HAE) (43), a parasitic disease endemic in certain regions where autologous liver transplantation offers a curative option for complex cases (44). Our analysis identified HAE as a current research hotspot (Cluster 2 in co-cited reference clustering), and Chinese institutions contribute to the literature in this specific area. While a detailed comparative analysis of Sino-US research on HAE is beyond the scope of this bibliometric study, the data suggest China's growing expertise in applying autotransplantation techniques to challenging regional diseases (45).

Italy, Japan, and the United Kingdom also have considerable influence in this field, with high publication volumes and citation rates reflecting their important roles in digestive system autologous transplantation research.

From an institutional perspective, the University of Minnesota System and the University of Minnesota Twin Cities are the institutions with the highest number of publications in this field, indicating their significant influence in digestive system autologous transplantation research. Analysis shows that these leading institutions not only have a high number of publications but also a high citation count of their research outcomes, reflecting widespread academic recognition and impact. Collaboration between domestic and international institutions shows that while domestic institutions collaborate closely, cross-national collaborations, especially between China and the United States, play a crucial role in advancing research in this field.

From a journal perspective, the American Journal of Transplantation and Transplantation Proceedings are the journals with the most publications, indicating their important position in digestive system autologous transplantation research. High-impact factor journals have significant authority and influence in this field, with Annals of Surgery and Transplantation being the most co-cited journals, further corroborating their core status in digestive system autologous transplantation research. These journals not only produce a high volume of research but also high-quality studies, representing the forefront and hotspots of this field.

Among all authors who have published literature on digestive system autologous transplantation, Bellin, Melena D., and Freeman, Martin L. are the most prolific. The research outcomes of these high-output authors have significantly contributed to the development of this field. Analysis indicates that these authors not only dominate in terms of publication volume but their research outcomes are also highly cited, reflecting their influence in the academic community. Further analysis of the author collaboration network shows that research in the field of digestive system autologous transplantation is highly collaborative, especially among prolific authors, which helps to advance the depth and breadth of research.

Through keyword analysis, we can identify the main hotspots in digestive system autologous transplantation research, including total pancreatectomy, islet autotransplantation, and chronic pancreatitis. The network diagram of high-frequency keywords shows close associations between different keywords, representing the main research directions and focal points in this field.

The co-occurrence analysis of keywords reveals several major clusters in this field of research: first, studies on autologous transplantation and related treatment methods; second, studies on pancreatic transplantation, its associated complications, and management; third, studies on surgical techniques and clinical outcomes. For instance, Cluster 1 (pink), which includes terms such as "autologous transplantation," "stem cell," "angiogenesis," "regeneration," "oxidative stress," "reperfusion injury," and "inflammation," points to fundamental biological processes and therapeutic strategies. "Stem cells," "angiogenesis," and "regeneration" reflect the ongoing research into enhancing organ repair and functional recovery post-transplantation. Concurrently, keywords like "oxidative stress," "reperfusion injury," and "inflammation" underscore critical challenges in transplantation, as these processes can significantly impact graft viability and patient outcomes; research in these areas aims to develop strategies for their mitigation. Cluster 2 (green), featuring "total pancreatectomy," "pain," "chronic pancreatitis," and "quality of life," directly relates to the primary clinical applications and patient-centered outcomes in pancreatic autotransplantation, particularly for debilitating chronic pancreatitis. Cluster 3 (purple), with keywords like "outcome," "surgery," "complication," "resection," and "tumor," encompasses the surgical aspects, oncological indications, and overall results of these complex procedures across different organs. Clusters 4 (yellow) and 5 (yellow), focusing on "TPIAT," "children," "insulin independence," "islet isolation," and "beta cell function," highlight the specialized area of islet autotransplantation, especially in pediatric populations and the efforts to preserve endocrine function. These research hotspots reflect the diversity and broad scope of research in digestive system autotransplantation.

Delving deeper into these hotspots, the prominence of total pancreatectomy with islet autotransplantation (TPIAT) for chronic pancreatitis is evident from its frequent appearance in keyword clusters (e.g., Cluster 2 and 4) and its focus in highly co-cited articles. TPIAT is significant because it offers a definitive solution for intractable pain in patients with end-stage chronic pancreatitis, a condition that severely impairs quality of life (46, 47). The simultaneous autotransplantation of islets aims to preserve endocrine function and prevent or mitigate brittle diabetes, a major drawback of total pancreatectomy alone (48). Current research in TPIAT, as suggested by keywords like "islet yield," "insulin independence," and "long-term outcome," focuses on optimizing patient selection, improving islet isolation and engraftment techniques to maximize insulin independence, and understanding long-term metabolic and quality-of-life outcomes, particularly in diverse populations including children. Challenges remain in consistently achieving high islet yields and durable insulin independence, as well as managing exocrine

pancreatic insufficiency post-surgery. Opportunities lie in refining islet processing, exploring adjuvant therapies to enhance islet survival (potentially linking to Cluster 1's themes of regeneration and cytoprotection), and standardizing postoperative management protocols.

The strong focus on "islet transplantation" itself, often in the context of TPIAT but also as a broader concept, underscores its critical role. Its significance lies in the potential to transform a lifealtering surgery (total pancreatectomy) into a more manageable condition by preserving endocrine function. The challenges revolve around the viability and functionality of transplanted islets, with research opportunities focused on enhancing isolation techniques, improving islet culture and preservation, and finding ways to protect islets from immediate post-transplant loss and ensure long-term survival and function (again, connecting to "beta cells," "inflammation," and "oxidative stress" from Cluster 1).

Furthermore, "surgical techniques" and "complications" are consistently highlighted keywords (e.g., Cluster 3). The complexity of digestive system autotransplantation means that refinements in surgical approaches-such as ex vivo resection, vascular reconstruction, and organ preservation-are paramount for success. The significance of advancing these techniques is direct: improved patient safety, reduced operative morbidity, and the ability to tackle increasingly complex cases, such as extensive tumors or severe anatomical distortions. Prominent challenges include managing complex vascular work, minimizing ischemia-reperfusion injury (a key term in Cluster 1), and reducing operative times. Opportunities for advancement include the adoption of minimally invasive techniques where feasible, enhanced intraoperative imaging and navigation, 3D modeling for preoperative planning, and further research into optimal organ preservation solutions (49). The high frequency of "complications" underscores the ongoing need to better understand, prevent, and manage adverse events such as bleeding, infection, thrombosis, and graft dysfunction. This presents opportunities for developing predictive risk models, standardizing perioperative care, and innovating strategies to mitigate common complications, thereby improving overall outcomes and patient recovery.

Further analysis shows that research hotspots vary across different stages. In the early stages, research primarily focused on basic research and initial clinical applications, such as technical improvements in islet transplantation and pancreatectomy and preliminary efficacy evaluations. In the mid-stages, research hotspots gradually shifted to clinical applications and management, such as managing complications post-pancreatic transplantation and improving patients' quality of life. In the later stages, as technology matured and clinical applications deepened, research focus shifted more toward long-term outcomes and large-scale clinical trials, such as long-term survival rates, preservation of islet function, and long-term improvement in patients' quality of life.

Although this study systematically analyzed literature in the field of digestive system autologous transplantation, there are still some limitations. For example, although the WoSCC database is highly authoritative, it may still miss some relevant literature not included in the database. Additionally, bibliometric methods primarily rely on quantitative indicators such as publication volume and citation counts, potentially overlooking some significant qualitative research outcomes.

Reflecting on the dynamism of the field, our analysis of citation bursts (Figure 9) revealed that 12 key references are currently in their peak citation period, suggesting their topics represent active and impactful areas of ongoing research within the last few years. While a comprehensive review of all recent highimpact articles is extensive, these burst references often point toward refinements in surgical techniques, long-term outcome assessments, and management of specific complications.

Future research could further combine qualitative analysis methods to deeply explore the intrinsic logic and development paths of research in digestive system autologous transplantation. Moreover, with the continuous emergence of new technologies and methods, research hotspots and directions in this field may change, necessitating continuous attention to these changes and timely adjustments in research strategies. For instance, future research might need to focus more on personalized treatment plans based on artificial intelligence and big data, as well as the application of new biomaterials and regenerative medicine in autologous transplantation. Additionally, emerging technologies such as 3D printing for surgical planning and the creation of patient-specific anatomical models or guides are beginning to show promise in complex surgeries and could influence future approaches in autotransplantation, although they may not yet be dominant themes in the retrospective bibliometric data up to 2024. The development of advanced imaging and intraoperative navigation systems also continues to enhance surgical precision.

Furthermore, interdisciplinary research and international cooperation will be crucial in advancing this field. By integrating knowledge and technologies from different disciplines and strengthening international collaboration, research outcomes can be accelerated for translation into clinical applications, thereby improving the efficacy of digestive system autologous transplantation and the quality of life for patients.

Conclusions

Our systematic analysis of the literature on autologous transplantation in the digestive system has uncovered the current state of research, key areas of interest, and evolving trends in this field. The findings demonstrate that the United States holds a dominant position, with significant contributions from countries like China. The primary research focuses are pancreatic transplantation, associated treatment methods, and clinical outcomes. To advance this field, future research should prioritize international collaboration and incorporate qualitative analysis techniques to deeply investigate the core aspects and developmental trajectories of related studies. This study not only maps the research landscape of autologous transplantation in the digestive system but also provides valuable references for future research. We hope this study will offer useful insights for researchers, thereby fostering the continued development and progress of autologous transplantation in the digestive system.

Data availability statement

The original contributions presented in the study are included in the article/Supplementary material, further inquiries can be directed to the corresponding authors.

Author contributions

AA: Writing – original draft, Writing – review & editing. TT: Writing – original draft, Writing – review & editing. AT: Data curation, Writing – review & editing. RZ: Data curation, Writing – review & editing. YS: Writing – review & editing. TA: Writing – review & editing.

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Conflict of interest

The authors declare that the research was conducted in the absence of any commercial or financial relationships that could be construed as a potential conflict of interest.

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Supplementary material

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