DOI: 10.35772/ghm.2020.01013

Effects of volume on outcome in hepatobiliary surgery: a review with guidelines proposal

Eloisa Franchi¹, Matteo Donadon^{1,2}, Guido Torzilli^{1,2,*}

¹Division of Hepatobiliary and General Surgery, Department of Surgery, Humanitas Clinical and Research Center – IRCCS, Rozzano, Milan, Italy; ²Department of Biomedical Science, Humanitas University, Pieve Emanuele, Milan, Italy.

Abstract: The positive relationship between volume and outcome in hepatobiliary surgery has been demonstrated for many years. As for other complex surgical procedures, both improved short- and long-term outcomes have been associated with a higher volume of procedures. However, whether the centralization of complex hepatobiliary procedures makes full sense because it should be associated with higher quality of care, as reported in the literature, precise criteria on what to centralize, where to centralize, and who should be entitled to perform complex procedures are still missing. Indeed, despite the generalized consensus on centralization in hepatobiliary surgery, this topic remains very complex because many determinants are involved in such a centralization process, of which some of them cannot be easily controlled. In the context of different health systems worldwide, such as national health systems and private insurance, there are different stakeholders that demand different needs: politicians, patients, surgeons, institutions and medical associations do not always have the same needs. Starting from a review of the literature on centralization in hepatobiliary surgery, we will propose some guidelines that, while not data-driven due to low evidence in the literature, will be based on good clinical practice.

Keywords: volume outcomes, hepatobiliary surgery, surgeon volume, centralization

Introduction

For years, the issue of centralization in liver surgery in specialist 'high-volume' hospitals has been prominent in the debate on improving quality in healthcare. It is well established that high volume, in general, means better outcome, and many studies have shown lower mortality and higher survival rates in high-volume versus low-volume centers (1-6). Indeed, in high-volume centers 90-day mortality rate is approximately 3%, with the morbidity rate around 30% (7-10). The factors involved seem to be many: better knowledge of the anatomy, more accurate selection of patients, refinements of surgical perioperative medicine techniques, as well as optimization of the management of postoperative complications (11-15).

The present review involves all available literature on the relationship between hospital or surgeon volume and postoperative mortality and survival in liver surgery suggesting some guidelines for management and creation of centralized departments.

Review of the literature

Table 1 details review of the literature regarding the relationship between outcome and volume in hepatobiliary surgery. Considering the rapid evolution of liver surgery, we have included articles published in the last 20 years in English. Moreover, we have included only those articles that have declassified hepatobiliary surgery from pancreatic surgery, which are usually considered together (16-45). As detailed, almost all the included articles supported a positive relationship between hospital volume and outcome indicating the validity of the union of high-volume and high-quality. In particular, in 2003, Dimick et al. (20) analyzed more than 2,000 hepatectomies performed in North America and found that those institutions that performed more than 20 resections per year had significantly lower mortality. Although the resulting cut off of 20 resections per year seems too inclusive, objectively the differences were substantial. However, in both groups the mean values outranged the benchmarks even of that period (6.3% vs. 15.5%). In 2009 a systematic review and in 2012 a meta-analysis confirmed a reduced mortality risk after liver surgery in high-volume centers (46,47).

Few of these articles, investigated how this relationship was mainly based on hospital or organization factors rather than on surgeon factors. In general, the positive relationship was evident both for the hospital and surgeon volumes. Even if this is reasonable, there

Author (Ref.)	Year	Patients	Importance of hospital volume	Importance of surgeon volume		
Begg CB, et al. (16)	1998	801	+	n/a		
Choti MA, et al. (17)	1998	606	+	n/a		
Glasgow RE, et al. (18)	1999	507	+	+		
Gordon TA, et al. (19)	1999	293	+	+		
Dimick JB, et al. (20)	2003	2,097	+	+		
Imamura H, et al. (21)	2003	1,056	+	+		
Fong Y, <i>et al.</i> (22)	2005	3,734	+	n/a		
Hollenbeck BK, et al. (23)	2007	3,630	+	n/a		
Eppsteiner RW, et al. (24)	2008	2,949	-	+		
McKay A, <i>et al.</i> (25)	2008	1,107	+	+		
Nathan H, et al. (26)	2009	6,871	+	-		
Stella M. (27)	2009	n/a	-	n/a		
Chamberlain RS, et al. (28)	2011	84	-	+		
Giuliante F, et al. (29)	2012	588	+	n/a		
Yasunaga H, et al. (30)	2012	18,046	+	n/a		
Viganò L, et al. (31)	2013	106	+	n/a		
Goetze TO, et al.* (32)	2014	487	+	n/a		
Ravaioli M, et al. (33)	2014	621	-	+		
Schneider EB, et al. (34)	2014	3,695	+	+		
Buettner S, et al. (15)	2014	9,874	+	n/a		
Aldrighetti L, et al.** (35)	2015	1,497	+	n/a		
Ejaz A, et al. (36)	2015	9,466	n/a	+		
Buettner S, et al. (37)	2016	5,075	+	+		
Gani F, et al. (38)	2016	27,813	+	n/a		
Botea F, et al. (39)	2017	3,016	+	+		
Chapman BC, et al. (40)	2017	12,757	+	+		
Idrees JJ, et al. (41)	2018	96,107	+	n/a		
Bouras AF, et al. ^{***} (42)	2019	46	-	n/a		
Chen Q, <i>et al.</i> (43)	2019	4,902	+	n/a		
Filmann N, et al. (44)	2019	110,332	+	+		
Chang CM, et al. (45)	2019	13,159	+	+		

Table 1	. Review of	f the liter	ature on	the r	relationship	between	outcome and	l volume	in l	nepatobiliar	y surge	ry
---------	-------------	-------------	----------	-------	--------------	---------	-------------	----------	------	--------------	---------	----

*Focus on gallbladder cancer; **Learning curve not hospital volume; ***Focus on laparoscopic liver surgery.

are confounding factors that are difficult to separate. In this sense, it is important to note that it is difficult to distinguish when high quality care in complex surgery is a consequence of reaching the plateau of a learning curve or when it is the consequence of a standard volume that is a minimum number of procedures per year. Besides, it is important to note that good outcomes in hepatobiliary surgery are also related to the quality of other hospital services, such as the anesthesiology service and the intensive care unit, which similarly to the surgeons have to reach the plateau of their learning curves. In this sense, further studies should be conducted to better characterize these two phenomena (*i.e.* learning curve versus minimum standard volume). Nathan H et al. (26) reported that the surgeon volume was not associated with in-hospital mortality, while Chang CM et al. (45) reported the combined effects of hospital and surgeon volume strongly influenced shortterm survival after hepatic resection. In this latter study, the prognosis was adjusted for several different factors such as indication for surgery, quality of the underlying chronic liver disease, and socio-economic status that were found to be important to be recorded and analyzed to strengthen the relationship between perioperative outcome and surgeon and/or hospital volume. Besides, Chang CM et al. (45) figured out that the combination

of high-volume surgeons in high-volume hospitals was associated with higher quality results, while the combination of high-volume surgeons in low-volume hospitals was not. Notably, in this study high-volume hospitals were those institutions performing more than 245 cases per year, while high-volume surgeons were those surgeons performing more than 59 cases per year. Notwithstanding these published studies, the definition of "high-volume center" remains to be elucidated. There is not an established cut-off of liver resections per year to perform (48).

Centralization of hepatobiliary surgery

The goal of centralization of hepatobiliary surgery is to provide optimal care of patients affected by hepatobiliary diseases within a given geographical area. This centralization passes through a complex process of assessment, development of dedicated policies, ongoing assurance and support from national government agencies, which should have the competence and authority to promote high quality care, good use of technical and technological tools, good allocation of human resources, and at the same time monitor, minimize and control the probability of unfortunate events. This process should be provided along a spacetime continuum that should warrant quality in all phases of the care of patients affected by hepatobiliary diseases.

These critical issues are very important in particular in liver surgery for several reasons. First, the definition of resectability is not standardized and wide variability is, in fact, observed among expert surgeons (49). Second, the complexity of liver surgery is difficult to be classified because several different types of resections requiring an extremely wide range of expertise can be performed. A standard distinction between major and minor hepatectomies is inadequate in the current era of modern liver surgery (50). Indeed, there are different technical solutions allowing parenchymal-sparing hepatectomies, much more complex than standard major hepatectomies, that remain in the shadow of the definition of minor hepatectomy. Yet, high quality centers should not be considered those centers performing a high proportion of major hepatectomies. In this sense, a new classification for minor hepatectomy that might help in better reporting minor but complex resections has been recently proposed (51). Third, postoperative morbidity and mortality rates have a limited validity to assess quality. Centers selecting only patients operable by performing small limited resections may have lower morbidity rates in comparison with centers routinely selecting patients operable by performing complex resections. Fourth, realistic cutoffs of mortality and morbidity rates after hepatectomy as a benchmark of quality should be defined to avoid the risk of denying the chance of care to those patients with higher complexity due to tumoral presentation or advanced age or because of severe comorbidities. Apart from the specificity of their indications for surgery, which requires being addressed by the local multidisciplinary teams (MDT), risk-adjusted metrics to compare outcomes among institutions are mandatory. Otherwise the risk of unfair comparisons will remain. In this sense, a benchmarking process has been started by merging the comprehensive complications risk (CCI) (52), liver failure occurrence, and morbidity and mortality classified according to the Clavien-Dindo classification (53). Last but not least, as recently pointed out by Aloia et al. (54) there are some downsides to the strategy of aiming at zero mortality rates after surgery such as the performance of innovative operations, which at least at the beginning are not compatible with perfection that might be strongly limited in the context of no-mortality. Therefore, the centralization process in hepatobiliary surgery should pass through the development and adoption of a new and modern common language for indications, resectability, terminology of resection, and good quality indicators.

Minimum hospital requirements in hepatobiliary surgery

To date, there are no specific published criteria that a

given hospital should have to perform hepatobiliary surgery. Most of the authors that have focused on this topic have reported their personal experiences, which anyway should be taken into consideration at least in the meantime of the reading out of some new studies with data. In 2016 a position paper published on behalf of the Italian Society of Surgery had the merit to feed up the debate and set some standards of reference (55). In Italy the current law about hospital standards is detailed by rule n. 70/2015, which divides hospitals into three levels (i.e. basic, I, and II levels). Accordingly, hepatobiliary surgery should be performed at least in level I hospitals or even better in level II hospitals, and the surgical team should be dedicated only to hepatobiliary and/or hepatobiliary and pancreatic procedures. This dedication should warrant a highquality standard.

Moreover, those high-quality hospitals, in which hepatobiliary surgery might be performed, should have the following departments: *i*) Department of Medical Oncology; *ii*) Department of Diagnostic Radiology, which should include some interventional radiologists dedicated to hepatobiliary diseases; *iii*) Department of Hepatology and/or of Internal Medicine with some internists dedicated to hepatobiliary diseases; *iv*) Department of Digestive Endoscopy; *v*) Intensive Care Unit; *vi*) Department of Pathology; *vii*) Department of Nuclear Medicine; and *viii*) Department of Radiation Oncology.

Even stating that the above-mentioned departments should be present in any high-quality hospital certified for hepatobiliary surgery, there might be a case of a given hospital that does not have some of the previous departments. In such a case, strong operative networks between that hospital and another institution should be activated to cover any deficiency. Similarly, in such a case of a given department of hepatobiliary surgery that does not provide liver transplantation another referral center in the same geographical area should be in the network to give consultation for liver transplantation. It should not be any more allowable that a patient with complex hepatobiliary disease hospitalized in a given hospital without the titles of performing diagnosis and/ or therapy for that specific disease do not provide the required network of care in the same geographical area.

Multidisciplinary team

Nowadays, it is mandatory to have MDT dedicated to patients affected by hepatobiliary diseases. MDT meetings provide the right global assessment of the patient both for diagnosis as well as for therapy. Any MDT meeting should include at least one member of the previous listed hospital departments with the aim to cover all the inherent aspects. Only physicians dedicated to liver diseases should take part to the MDT meeting, which should be scheduled based on the case-load but in general once per week. A written report of the MDT should be provided for each patient with the signature of all those members that have contributed to the discussion. It is important to note that the correct functioning of the MDT meeting relies on the proper union between the scientific evidence and the local experience in the diagnosis and cure of a given hepatobiliary disease. A MDT well balanced among specialties represented, and authoritative in all its specialists, provides better patient management resulting in better short- and long-term outcomes (56-59).

Hospital volume versus surgeon volume

Ideally, hospital volume and surgeon volume should match while in the real world this is not always warranted. In hepatobiliary surgery, the relative importance of hospital versus surgeon volume is very important because both short- and long-term outcomes are dependent on hospital factors, such as the presence of intensive care unit, and surgeon factors, such as the operative technique. Nathan H et al. (26) showed that the protective effect of hospital hepatic resection volume persisted after case-mix adjustment for competing risk factors, while that was not the case considering the surgeon hepatic resection volume. Indeed, high- and low-volume surgeons had comparable in-hospital mortality rates after hepatectomy (26). There are also other factors inherent in the hospital organization which were not considered and may have biased Nathan et al. conclusions: i.e. an active MDT meeting discussing each patient as above stated, which was not considered by them and by many other authors as well.

Learning curve or standard volume?

Center volume, surgeon volume, and surgeon experience all appear to impact success rates in liver surgery. A better understanding of how these factors interact to influence outcomes could help to develop specific healthcare strategies for the improvement of the quality of care in patients with hepatobiliary diseases. As said before, it is difficult to distinguish if good outcomes in hepatobiliary surgery are more dependent on the learning curve or to a minimum standard volume. A possible strategy to overcome this infertile dualism might be the introduction of certification for hepatobiliary surgeons. Far from the idea of more bureaucracy, this strategy might include analysis of the training with emphasis on the schools of surgery, and mentors that a given surgeon might have trained under during his or her career to be entitled in performing complex hepatobiliary surgery. As recently pointed out by some authors, this was found to be a good strategy in the field of pancreatic surgery and might work also in

other fields of surgery (31,59). Besides, it might be the way to reinforce the importance of schools of surgery, which are those named to train young surgeons.

Toward certified hepatobiliary surgeons

A strategy to overcome the difficulty in decoding the dualism hospital volume - surgeon volume might be the introduction of certification provided by a national board of specialists. This board should be an independent, non-profit organization founded for the purpose of certifying surgeons who have met a defined standard of education, training and knowledge. Moreover, this board might work in defining the minimum standard of care in hepatobiliary surgery on an individual basis and might analyze the applicant's training and operative experience as well as his/her professionalism and ethics. Upon successful completion of these analyses, the surgeon might become certified in hepatobiliary surgery. This certification might serve as a prerequisite of good practice in hepatobiliary surgery, which together with the above reported minimum hospital requirements in hepatobiliary surgery, both as a single institution or as an established network between different institutions, might be warranted for highquality care - independently by a number of procedures. Notably once certified, the hepatobiliary surgeon should undergo a process of maintenance of certification (every 5-10 years) with the aim of demonstrating ongoing professionalism and commitment to continuing medical education in the field of hepatobiliary surgery.

Conclusions

In conclusion, volume and outcome data in hepatobiliary surgery are intrinsically associated with some limitations. The published studies are mostly observational, and retrospective. Besides, the centralization process requires preparatory and preliminary agreements among experts about the development and adoption of new and modern common language for indications, resectability, terminology of resection, and good quality indicators. Without these agreements, hospital as well as surgeon volume act as proxy measures for technical and nontechnical skills. However, such a centralization process remains very important to offer better care for patients suffering from complex hepatobiliary disease.

References

- Luft HS BJ, Enthoven AC. Should operations be regionalized? The empirical relation between surgical volume and mortality. N Engl J Med. 1979; 301:1364-1369.
- 2. Luft HS. The relation between surgical volume and mortality: an exploration of causal factors and alternative models. Med Care. 1980; 18:940-959.
- 3. GordonTA, Burleyson GP, Tielsch JM, Cameron JL. The

effects of regionalization on cost and outcome for one general high-risk surgical procedure. Ann Surg. 1995; 221:43-49.

- Birkmeyer JD, Siewers AE, Finlayson EV, Stukel TA, Lucas FL, Batista I, Welch HG, Wennberg DE. Hospital volume and surgical mortality in the United States. N Engl J Med. 2002; 346:1128-1137.
- Finlayson EV, Goodney PP, Birkmeyer JD. Hospital volume and operative mortality in cancer surgery: a national study. Arch Surg. 2003; 138:721-725; discussion 726.
- Birkmeyer JD, StukelTA, Siewers AE, Goodney PP, Wennberg DE, Lucas FL. Surgeon volume and operative mortality in the United States. N Engl J Med. 2003; 349:2117-2127.
- Dimick JB, Pronovost PJ, Cowan JA Jr, Lipsett PA. Postoperative complication rates after hepatic resection in Maryland hospitals. Arch Surg. 2003; 138:41-46.
- Dimick JB, Wainess RM, Cowan JA, Upchurch GR Jr, Knol JA, Colletti LM. National trends in the use and outcomes of hepatic resection. J Am Coll Surg. 2004; 199:31-38.
- Finks JF, Osborne NH, Birkmeyer JD. Trends in hospital volume and operative mortality for high-risk surgery. N Engl J Med. 2011; 364:2128-2137.
- Liu JH, Zingmond DS, McGory ML, SooHoo NF, Ettner SL, Brook RH, Ko CY. Disparities in the utilization of high-volume hospitals for complex surgery. JAMA. 2006; 296:1973-1980.
- Pearse RM, Moreno RP, Bauer P, Pelosi P, Metnitz P, Spies C, Vallet B, Vincent JL, Hoeft A, Rhodes A; European Surgical Outcomes Study (EuSOS) group for the Trials groups of the European Society of Intensive Care Medicine and the European Society of Anaesthesiology. Mortality after surgery in Europe: a 7 day cohort study. Lancet. 2012; 380:1059-1065.
- Bauer H, Honselmann KC. Minimum Volume Standards in Surgery - Are We There Yet? Visc Med. 2017; 33:106-116.
- Morche J, Mathes T, Pieper D. Relationship between surgeon volume and outcomes: a systematic review of systematic reviews. Syst Rev. 2016; 5:204.
- Spolverato G, Ejaz A, Hyder O, Kim Y, Pawlik TM. Failure to rescue as a source of variation in hospital mortality after hepatic surgery. Br J Surg. 2014; 101:836-846.
- Buettner S, Gani F, Amini N, Spolverato G, Kim Y, Kilic A, Wagner D, Pawlik TM. The relative effect of hospital and surgeon volume on failure to rescue among patients undergoing liver resection for cancer. Surgery. 2016; 159:1004-1012.
- Begg CB, Cramer LD, Hoskins WJ, Brennan MF. Impact of hospital volume on operative mortality for major cancer surgery. JAMA. 1998; 280:1747-1751.
- Choti MA, Bowman HM, Pitt HA, Sosa JA, Sitzmann JV, Cameron JL, Gordon TA. Should hepatic resections be performed at high-volume referral centers? J Gastrointest Surg. 1998; 2:11-20.
- Glasgow RE, Showstack JA, Katz PP, Corvera CU, Warren RS, Mulvihill SJ. The relationship between hospital volume and outcomes of hepatic resection for hepatocellular carcinoma. Arch Surg. 1999; 134:30-35.
- Gordon TA, Bowman HM, Bass EB, Lillemoe KD, Yeo CJ, Heitmiller RF, Choti MA, Burleyson GP, Hsieh G, Cameron JL. Complex gastrointestinal surgery: impact of

provider experience on clinical and economic outcomes. J Am Coll Surg. 1999; 189:46-56.

- 20. Dimick JB, Cowan JA Jr, Knol JA, Upchurch GR Jr. Hepatic resection in the United States: indications, outcomes, and hospital procedural volumes from a nationally representative database. Arch Surg. 2003; 138:185-191.
- Imamura H, Seyama Y, Kokudo N, Maema A, Sugawara Y, Sano K, Takayama T, Makuuchi M. One thousand fifty-six hepatectomies without mortality in 8 years. Arch Surg. 2003; 138:1198-1206.
- 22. Fong Y, Gonen M, Rubin D, Radzyner M, Brennan MF. Long-term survival is superior after resection for cancer in high-volume centers. Ann Surg. 2005; 242:540-547.
- Hollenbeck BK, Dunn RL, Miller DC, Daignault S, Taub DA, Wei JT. Volume-based referral for cancer surgery: Informing the debate. J Clin Oncol. 2007; 25:91-96.
- Eppsteiner RW, Csikesz NG, Simons JP, Tseng JF, Shah SA. High volume and outcome after liver resection: surgeon or center? J Gastrointest Surg. 2008; 12:1709-1716.
- McKay A, You I, Bigam D, Lafreniere R, Sutherland F, Ghali W, Dixon E. Impact of surgeon training on outcomes after resective hepatic surgery. Ann Surg Oncol. 2008; 15:1348-1355.
- Nathan H, Cameron JL, Choti MA, Schulick RD, Pawlik TM. The volume-outcomes effect in hepato-pancreatobiliary surgery: hospital versus surgeon contributions and specificity of the relationship. J Am Coll Surg. 2009; 208:528-538.
- Stella M. Safety and feasibility of liver resection at lowvolume institutions. Surgery. 2009; 145:575.
- 28. Chamberlain RS, Klaassen Z, Paragi PR. Complex hepatobiliary surgery in the community setting: is it safe and feasible? Am J Surg. 2011; 202:273-280.
- Giuliante F, Ardito F, Pinna AD, *et al.* Liver resection for hepatocellular carcinoma ≤ 3 cm: results of an Italian multicenter study on 588 patients. J Am Coll Surg. 2012; 215:244-254.
- Yasunaga H, Horiguchi H, Matsuda S, Fushimi K, Hashimoto H, Ohe K, Kokudo N. Relationship between hospital volume and operative mortality for liver resection: data from the Japanese Diagnosis Procedure Combination database. Hepatol Res. 2012; 42:1073-1080.
- Viganò L, Langella S, Ferrero A, Russolillo N, Sperti E, Capussotti L. Colorectal cancer with synchronous resectable liver metastases: monocentric management in a hepatobiliary referral center improves survival outcomes. Ann Surg Oncol. 2013; 20:938-945.
- Goetze TO, Paolucci V. Influence of high- and lowvolume liver surgery in gallbladder carcinoma. World J Gastroenterol. 2014; 20:18445-18451.
- 33. Ravaioli M, Pinna AD, Francioni G, Montorsi M, Veneroni L, Grazi GL, Palini GM, Gavazzi F, Stacchini G, Ridolfi C, Serenari M, Zerbi A. A partnership model between high- And low-volume hospitals to improve results in hepatobiliary pancreatic surgery. Ann Surg. 2014; 260:871-877.
- 34. Schneider EB, Ejaz A, Spolverato G, Hirose K, Makary MA, Wolfgang CL, Ahuja N, Weiss M, Pawlik TM. Hospital volume and patient outcomes in hepatopancreatico-biliary surgery: is assessing differences in mortality enough? J Gastrointest Surg. 2014; 18:2105-2115.
- 35. Aldrighetti L, Belli G, Boni L, Cillo U, Ettorre G, De

Carlis L, Pinna A, Casciola L, Calise F; Italian Group of Minimally Invasive Liver Surgery (I GO MILS). Italian experience in minimally invasive liver surgery: a national survey. Updates Surg. 2015; 67:129-140.

- 36. Ejaz A, Spolverato G, Kim Y, Wolfgang CL, Hirose K, Weiss M, Makary MA, Pawlik TM. The impact of resident involvement on surgical outcomes among patients undergoing hepatic and pancreatic resections. Surgery. 2015; 158:323-330.
- Buettner S, Gani F, Amini N, Spolverato G, Kim Y, Kilic A, Wagner D, Pawlik TM. The relative effect of hospital and surgeon volume on failure to rescue among patients undergoing liver resection for cancer. Surgery. 2016; 159:1004-1012.
- Gani F, Pawlik TM. Assessing the costs associated with volume-based referral for hepatic surgery. J Gastrointest Surg. 2016; 20:945-952.
- Botea F, Ionescu M, Braşoveanu V, *et al.* Liver resections in a high-volume center: Form standard procedures to extreme surgery and ultrasound-guided resections. Chirurgia (Bucur). 2017; 112:259-277.
- 40. Chapman BC, Paniccia A, Hosokawa PW, Henderson WG, Overbey DM, Messersmith W, McCarter MD, Gleisner A, Edil BH, Schulick RD, Gajdos C. Impact of facility type and surgical volume on 10-year survival in patients undergoing hepatic resection for hepatocellular carcinoma. J Am Coll Surg. 2017; 224:362-372.
- 41. Idrees JJ, Kimbrough CW, Rosinski BF, Schmidt C, Dillhoff ME, Beal EW, Bagante F, Merath K, Chen Q, Cloyd JM, Ellison EC, Pawlik TM. The cost of failure: assessing the cost-effectiveness of rescuing patients from major complications after liver resection using the national inpatient sample. J Gastrointest Surg. 2018; 22:1688-1696.
- Bouras AF, Liddo G, Marx-Deseure A, Leroy A, Decanter G. Accessible laparoscopic liver resection performed in low volume centers: Is it time for democratization? J Visc Surg. 2020; 157:193-197.
- 43. Chen Q, Olsen G, Bagante F, Merath K, Idrees JJ, Akgul O, Cloyd J, Dillhoff M, White S, Pawlik TM. Procedure-specific volume and nurse-to-patient ratio: implications for failure to rescue patients following liver surgery. World J Surg. 2019; 43:910-919.
- Filmann N, Walter D, Schadde E, Bruns C, Keck T, Lang H, Oldhafer K, Schlitt HJ, Schön MR, Herrmann E, Bechstein WO, Schnitzbauer AA. Mortality after liver surgery in Germany. Br J Surg. 2019; 106:1523-1529.
- 45. Chang CM, Yin WY, Wei CK, Lee CH, Lee CC. The combined effects of hospital and surgeon volume on short-term survival after hepatic resection in a population-based study. PLoS One. 2014; 9:e86444.
- Garcea G, Breukink SO, Marlow NE, Maddern GJ, Barraclough B, Collier NA, Dickinson IC, Fawcett J, Graham JC. A systematic review of the impact of volume of hepatic surgery on patient outcome. Surgery. 2009; 145:467-475.
- Tol JA, van Gulik TM, Busch OR, Gouma DJ. Centralization of highly complex low-volume procedures in upper gastrointestinal surgery. A summary of systematic reviews and meta- analyses. Dig Surg. 2012; 29:374-383.
- Gruen RL, Pitt V, Green S, Parkhill A, Campbell D, Jolley D. The effect of provider case volume on cancer mortality: systematic review and meta-analysis. CA

Cancer J Clin. 2009; 59:192-211.

- 49. Folprecht G, Gruenberger T, Bechstein WO, *et al.* Tumour response and secondary resectability of colorectal liver metastases following neoadjuvant chemotherapy with cetuximab: the CELIM randomised phase 2 trial. Lancet Oncol. 2010; 11:38-47.
- Pang YY. The Brisbane 2000 terminology of liver anatomy and resections. HPB 2000; 2:333-39. HPB (Oxford). 2002; 4:99; author reply 99-100.
- 51. Viganò L, Torzilli G, Troisi R, *et al.* Minor hepatectomies: focusing a blurred picture: analysis of the outcome of 4471 open resections in patients without cirrhosis. Ann Surg. 2019; 270:842-851.
- 52. Slankamenac K, Graf R, Barkun J, Puhan MA, Clavien PA. The comprehensive complication index: a novel continuous scale to measure surgical morbidity. Ann Surg. 2013; 258: 1-7
- 53. Rössler F, Sapisochin G, Song G, *et al.* Defining benchmarks for major liver surgery: a multicenter analysis of 5202 living liver donors. Ann Surg. 2016; 264:492-500.
- 54. Aloia TA. Should zero harm be our goal? Ann Surg. 2020; 271:33-36.
- 55. Torzilli G, Viganò L, Giuliante F, Pinna AD. Liver surgery in Italy. Criteria to identify the hospital units and the tertiary referral centers entitled to perform it. Updates Surg. 2016; 68:135-142.
- 56. Yopp AC, Mansour JC, Beg MS, Arenas J, Trimmer C, Reddick M, Pedrosa I, Khatri G, Yakoo T, Meyer JJ, Shaw J, Marrero JA, Singal AG. Establishment of a multidisciplinary hepatocellular carcinoma clinic is associated with improved clinical outcome. Ann Surg Oncol. 2014; 21:1287-1295.
- 57. Gashin L, Tapper E, Babalola A, Lai KC, Miksad R, Malik R, Cohen E. Determinants and outcomes of adherence to recommendations from a multidisciplinary tumour conference for hepatocellular carcinoma. HPB (Oxford). 2014; 16:1009-1015.
- 58. Viganò L, Pedicini V, Comito T, Carnaghi C, Costa G, Poretti D, Franzese C, Personeni N, Del Fabbro D, Rimassa L, Scorsetti M, Santoro A, Solbiati L, Torzilli G. Aggressive and multidisciplinary local approach to iterative recurrences of colorectal liver metastases. World J Surg. 2018; 42:2651-2659.
- 59. Capretti G, Balzano G, Gianotti L, Stella M, Ferrari G, Baccari P, Zuliani W, Braga M, Zerbi A. Management and outcomes of pancreatic resections performed in highvolume referral and low-volume community hospitals lead by surgeons who shared the same mentor: the importance of training. Dig Surg. 2018; 35:42-48.

Received March 12, 2020; Revised July 29, 2020; Accepted August 11, 2020.

Released online in J-STAGE as advance publication September 4, 2020.

*Address correspondence to:

Guido Torzilli, Division of Hepatobiliary and General Surgery, Department of Surgery, Humanitas Clinical and Research Center – IRCCS, Via Manzoni 56, 20089, Rozzano, Milan, Italy.

E-mail: guido.torzilli@hunimed.eu