Global experience and progress in GreenLight-XPS 180-Watt photoselective vaporization of the prostate

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Abstract
Purpose To evaluate changes in global perioperative data of GreenLight-XPS 180-Watt photo-selective vaporization of the prostate (GL-XPS) of the Global Greenlight Group (GGG) database.
Methods 3441 men, who underwent GL-XPS for symptomatic BPH between 2011 and 2019 at seven high volume international centers, were included. Primary outcome measurements were operative time (OT; min), effective laser time (LT; min of OT), as well as intraoperative and postoperative adverse events (AEs), all analyzed by year of surgery (2011–2019) and prostate volume (PV) group (< 80 ml vs. 80-150 ml vs. > 150 ml).
Results The median age was 70 years (interquartile range 64–77), the median PV was 64 ml (IQR 47–90). The OT and LT slightly increased but stayed highly efficient all in all. Median OT was 60 min (IQR 45–83) and LT was 33 min (IQR 23–46). Median energy use was 253 kJ (IQR 170–375) with an energy density of 3.94 kJ/ml (IQR 2.94–5.02). The relative probability of perioperative AEs decreased by 17% each year (p < 0.001). The relative probability of perioperative transfusion dropped significantly from 2% in 2011 to 0% in 2019 (p = 0.007). The early postoperative complications (within 30 days after surgery) decreased significantly from 48.8% (n = 106) in 2011 to 24.7% (n = 20) in 2019 (p > 0.001).
Conclusion These findings from the GGG demonstrate significant improvement secondary to growing experience with GL-XPS between 2011 and 2019 in intraoperative AEs, including transfusions, and postoperative AEs. While staying highly efficient in OT and LT of GL-XPS within a 9-year period of experience.

Keywords Global experience · Greenlight · BPH · GL-XPS · Global Greenlight Group

Introduction
Following the failure of conservative medical management to adequately address lower urinary tract symptoms (LUTS) and elevated post-void urine volume secondary to benign prostate hyperplasia (BPH), surgical therapy is often recommended. Transurethral resection of the prostate (TURP) remains the current surgical gold standard for prostates between 30 and 80 ml [1] but this procedure is associated with several perioperative adverse events (AEs) such as bleeding [2], especially in larger prostates [3]. GreenLight-XPS 180-Watt photoselective vaporization of the prostate (GL-XPS) has been added to the international guidelines as a safe surgical method with decreased morbidity especially in prostates > 100 ml and for patients requiring therapeutic anticoagulation therapy [1–5].

In 2014, the GOLIATH study confirmed non-inferiority of GL-XPS compared to TURP regarding IPSS-QoL, Qmax, and AEs [6]. Since then, with growing global experience, the surgical technique and application of GL-XPS have certainly evolved. We have previously shown that growing experience with GL-XPS at a single high-volume center is associated with improved OT and LT over a 5-year period [7]. However, there are currently no studies that have evaluated the global...
progress of GL-XPS in respect to effectiveness, efficacy, and safety.

As such, this study sought to evaluate the perioperative progress of GL-XPS secondary to growing global experience using data from the Global Greenlight Group (GGG) database which represents multiple, international high-volume centers. The primary outcomes were operative and laser time (OT, LT; min), as well as intraoperative and postoperative AEs, all analyzed by prostate volume (PV) (ml) and by the year of surgery.

**Patients and methods**

**Patients and study design**

This retrospective analysis of the GGG database included 3441 men who underwent GL-XPS (Boston Scientific, Minnetonka, MN, USA) for symptomatic BPH between 2011 and 2019 from one of eight experienced surgeons at seven international high-volume centers in Germany, Canada, France, Italy, Mexico, Brazil, and Argentina [8]. Our study
design was visualized in Fig. 1. High Data quality/density was ensured using objective variables and a close cooperation between the individual centers of the Global Greenlight Group.

**Study outcomes**

Primary outcome measurements were OT, LT, perioperative AEs, and perioperative transfusions analyzed by year of surgery (2011–2019) and by PV (PV1: < 80 ml vs. PV2: 80–150 ml vs. PV3: > 150 ml) with respect to effectiveness, efficacy, and safety with a postoperative follow-up of 12 months.

Postoperative follow-up included International Prostate Symptom Score–Quality of Life (IPSS–QoL), early postoperative AE (< 30 days, subcategorized in low-grade (Clavien-Dindo grade I–II) and high-grade (need of an intervention; Clavien-Dindo grade > IIIa–IV), hospital length of stay (days), catheterization time (days), and postoperative re-catheterization rate.

**Surgical procedures**

All eight GL-XPS surgeons were fellowship-trained urologists equally experienced in performing GL-XPS regularly. GL-XPS was performed using the XPS-180 W system and MoXy™ laser fiber. Procedures were performed in accordance to published international guidelines [2, 9]. Procedures were performed under general or spinal anesthesia. Prophylactic antibiotic treatment, as well as antithrombotic therapy were administered according to local guidelines.

**Statistical analysis**

Non-normally distributed data are presented as a median (IQR) or as a percentage. Categorical variables were presented by relative frequencies. Linear regression analysis and logistic regression analysis were conducted for the data set to visualize the trend of the years of surgery. Multivariable logistic regression analysis was performed to detect influencing factors. Statistical significance was set at a two-sided p < 0.05. Data were analyzed with SPSS, version 27 (Chicago, Illinois, USA).

**Results**

**Preoperative baseline results**

Patient baseline characteristics of the 3441 men are summarized in Table 1. The median age was 70 years (interquartile range IQR 64–77) with a median PV of 64 ml (IQR 47–90) and a median PSA level of 3.27 ng/dl (IQR 1.76–5.88). Between 2011 and 2019, there was a significant increase in PV treated (p = 0.02) by 0.71 ml each year. In 2011, there were 28.7% of patients with PV2, 3% PV3 and in 2019 it raised to 43.6% in PV2 and 4% in PV3. The median IPSS/QoL was 23/4 (IQR 19–27/3–5).

The median post-void residual volume was 120 ml (IQR 30–273) and showed a significant decrease between the years of surgery (p < 0.001). Additionally, there was a significant increase in patients with a median lobe (p < 0.001) rising from 36.2% in 2011 to 64% in 2019.

With regards to anticoagulation status, 35% (n = 1176) of men undergoing surgery were prescribed an oral anticoagulation such as acetylsalicylic acid or coumadin due to co-existing medical conditions. Overall, there was a significant increase of patients on general medication (such as anticoagulation, anti-diabetics or pain killers) between 2011 and 2019 (p < 0.001).

**Intraoperative baseline results**

The analysis showed a significant increase in OT (p < 0.001) and LT (p < 0.001)—see Fig. 2a, b. In subgroup analyses by PV, the OT only showed a significant increase in the PV2 group (81–150 ml) while the LT showed a significant increase in PV1-3. Although the prostate glands in PV3 were significantly larger (up to 330 ml), the LT only increased by approximately 3 min per year (p = 0.011), while the LT in PV1 (p < 0.001) and PV2 (p < 0.001) only increased by approximately 2 min per year.

The applied energy (median 253 kJ, IQR 170–375, PV1 p = 0.338, PV2 p = 0.725, PV3 p = 0.931) stayed constant with no change in energy density (median 3.94 kJ/ml, IQR 2.94–5.02, PV1 p = 0.344, PV2 p = 0.993, PV3 p = 0.192) in the observation period between the PV 1–3 while the energy used (p = 0.038) showed a significant increase overall.

The median LT/OT ratio stayed highly efficient over 50% for PV1-3. It was significantly improved between 2011 and 2019 for PV1 (p < 0.001) and PV2 (p < 0.001) up to over 60% in 2017 and 2018.

Most patients were treated with one fiber (75.6%). The LT per PV (median 0.5 min/ml, IQR 0.38–0.66) showed a steady increase with 0.025 min/ml (p < 0.001) each year. The most obvious increase was observed in PV1 with an increase of 0.029 min/ml each year (p < 0.001).

Vice versa the applied laser energy per LT (median 8.45 kJ/min, IQR 6.72–9.44) showed a steady decrease (p < 0.001) between 2011 and 2019. Moreover, there was a significant decrease in PV1-3 by approximately 0.3 kJ/min each year as shown in Fig. 2e.

With regard to perioperative AEs, conversion to TURP, which was the most frequent AE observed, decreased from 16% (n = 24) in 2011 to 9.4% (n = 6) in 2019 (p < 0.001) as shown in Fig. 2g. Likewise, perioperative transfusions dropped from 2% in 2011 to 0% in 2019 (p = 0.007).
| Table 1  | Patients demographics between 2011 and 2019 (linear regression or binar logistic regression) |
|----------|------------------------------------------------------------------------------------------|
| n = max. 3441 | Baseline | 2011 | 2012 | 2013 | 2014 |
| | Median/n (IQR/%) | Median/n | IQR/% | Median/n | IQR/% | Median/n | IQR/% | Median/n | IQR/% | Median/n | IQR/% |
| Age, years | 70 (64–77) | 70 | 65–77 | 71 | 64–77 | 70 | 63–77 | 70 | 64–76 |
| PSA, ng/dl | 3.27 (1.76–5.88) | 3.76 | 2–7 | 2.84 | 1.60–5.48 | 3.55 | 2–5.6 | 3.2 | 1.9–5.4 |
| TRUS volume, ml | 64 (47–90) | 61 | 49–89 | 62.00 | 43–88 | 63 | 45–83 | 63 | 48–90 |
| IPSS baseline | 23 (19–27) | 23 | 19–27 | 22.50 | 18–29 | 22 | 19–28 | 23 | 19–27 |
| QoL baseline | 4 (3–5) | 5 | 4–5 | 5.00 | 4–6 | 5 | 4–5 | 5 | 4–5 |
| Qmax baseline, ml/s | 6 (4–8) | 6 | 4–8 | 5.5 | 3–7 | 6 | 4–7 | 5.5 | 4–8 |
| PVR, ml | 120 (30–273) | 175 | 80–418 | 151.5 | 66–400 | 168 | 65.5–314 | 126.5 | 50–333 |
| Medication use | 729 (60.6%) | 75 | 52.8% | 101 | 54.6% | 88 | 53.7% | 88 | 55.7% |
| Anticoagulation use | 1176 (35%) | 87 | 39.7% | 115 | 33.8% | 113 | 31.6% | 173 | 34.7% |
| α-blocker use | 2661 (79.9%) | 182 | 83.9% | 280 | 83.3% | 288 | 81.4% | 390 | 80.1% |
| 5α-reductase inhibitor use | 1424 (42.6%) | 118 | 54.1% | 179 | 53.1% | 159 | 49.9% | 176 | 36.1% |
| Median Lobe | 532 (36.2%) | 54 | 36.2% | 77 | 36.0% | 57 | 26.4% | 57 | 29.2% |
| ASA | 428 (22%) | 28 | 18.9% | 59 | 25.4% | 37 | 16.5% | 32 | 16.9% |
| 996 (51.2%) | 83 | 56.1% | 108 | 53.1% | 106 | 47.3% | 88 | 46.6% |
| 504 (25.9%) | 35 | 23.6% | 62 | 26.7% | 78 | 34.8% | 66 | 34.9% |
| 18 (0.9%) | 12 | 7.1% | 5 | 3.3% | 3 | 1.5% | 3 | 1.6% |
| Retention, preoperative | 342 (35.8%) | 39 | 47.6% | 45 | 36.3% | 53 | 42.4% | 54 | 43.9% |
| Foley in place, preoperative | 841 (27.7%) | 69 | 31.4% | 90 | 28.6% | 94 | 29.8% | 104 | 24.8% |

| n = max. 3441 | Baseline | 2015 | 2016 | 2017 | 2018 | 2019 |
| | Median/n (IQR/%) | Median/n | IQR/% | Median/n | IQR/% | Median/n | IQR/% | Median/n | IQR/% |
| Age, years | 70 (64–77) | 69 | 64–76 | 70 | 64–76 | 71 | 65–79 | 70 | 64–77 |
| PSA, ng/dl | 3.27 (1.76–5.88) | 3.2 | 1.7–5.2 | 3 | 1.52–5.76 | 3.15 | 1.62–5.7 | 3.61 | 1.91–7.33 |
| TRUS volume, ml | 64 (47–90) | 65 | 47–88 | 61 | 46–85 | 64.5 | 47–90 | 70.5 | 50–96 |
| IPSS baseline | 23 (19–27) | 23 | 19–27 | 22 | 19–27 | 22 | 19–27 | 22 | 17–27 |
| QoL baseline | 4 (3–5) | 5 | 4–5 | 4 | 3–5 | 3 | 0–5 | 4 | 2–5 |
| Qmax baseline, ml/s | 6 (4–8) | 6 | 4–9 | 6 | 4–8 | 6 | 4–8 | 6 | 4–7 |
| PVR, ml | 120 (30–273) | 112.5 | 20–230 | 100 | 0–200 | 99.5 | 0–200 | 67.5 | 0–191.5 |
| Medication use | 729 (60.6%) | 71 | 58.7% | 74 | 63.8% | 78 | 75.0% | 86 | 78.9% |
| Anticoagulation use | 1176 (35%) | 194 | 34.5% | 219 | 36.9% | 129 | 35.1% | 115 | 36.2% |
| α-blocker use | 2661 (79.9%) | 429 | 77.3% | 455 | 76.9% | 304 | 84.0% | 240 | 75.2% |
| 5α-reductase inhibitor use | 1424 (42.6%) | 209 | 37.6% | 219 | 37.0% | 173 | 47.0% | 133 | 41.7% |
| Median Lobe | 532 (36.2%) | 59 | 35.8% | 48 | 29.4% | 56 | 37.6% | 60 | 51.3% |
| ASA | 428 (22%) | 56 | 17.8% | 86 | 20.9% | 55 | 29.4% | 51 | 36.4% |
| 996 (51.2%) | 174 | 55.2% | 224 | 54.5% | 94 | 50.3% | 71 | 50.7% |
| 504 (25.9%) | 84 | 26.7% | 98 | 23.8% | 36 | 19.3% | 18 | 12.9% |
| 18 (0.9%) | 1 | 0.3% | 3 | 0.7% | 2 | 1.1% | 0 | 0.0% |
| Retention, preoperative | 342 (35.8%) | 35 | 32.4% | 42 | 35.3% | 30 | 25.2% | 24 | 26.7% |
| Foley in place, preoperative | 841 (27.7%) | 129 | 25.4% | 138 | 25.7% | 104 | 31.8% | 96 | 31.8% |

| n = max. 3441 | Baseline | F | Coefficient variable | Coefficient constant | R² adjusted | R² | p |
| | Median/n (IQR/%) | | | | | | |
| Age, years | 70 (64–77) | 70 | 0.778 | 0.778 | 0.778 | 0.778 | 0.778 |

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The subanalysis of PV1-3 showed no reduction in intraoperative AEs compared to PV3 (\(p = 0.967\)).

**Postoperative baseline results**

The length of hospital stay remained consistent with a median of two days (\(p = 0.553\)) and a median postoperative catheterization time of 1 day with a significantly shorter range in PV3 (\(p = 0.043\)). The need for re-catherization stayed constant across the years (\(p = 0.433, 5.9\%\)).

Overall, the preoperative IPSS/QoL (median 23/4, IQR 19–27/3–5) showed a reduction in symptomatic BPO after 3 months (median 6/1, IQR 4–9–1–2, \(p > 0.001\)) and remained stable at both 6 (median 5/1, IQR 4–9–0–1) and 12 months (median 4/1; IQR 2–7/0–1). Between 2011 and 2019 the preoperative QoL showed a significant decrease by \(-0.188\) each year (\(p < 0.001\)). Moreover, there was a significant reduction of QoL after 12 months of surgery (\(p = 0.005\)).

The median postvoid residual volume was reduced from 120 ml (IQR 30–273) to 30 ml (IQR 0–75, \(p < 0.001\)) post-operatively, without any changes over the years, and a stable result in our follow-up time up to 12 months after surgery. The median Qmax improved from preoperative 6 ml/s (IQR 4–8) to 3 months postoperative 20 ml/s (IQR 16–22, \(p < 0.001\)). The outcome stayed stable after 6 and 12 months and between 2011 and 2019.

**Postoperative adverse events**

Overall postoperative AEs (Clavien-Dindo classification low-grade and high-grade) were significantly reduced from 2011 with \(n = 106\) (48.8\%) to 2019 with \(n = 20\) (24.7\%) (\(p < 0.001\)). The sub-analysis of PV1-3 showed a significant decrease in postoperative AE especially for prostates from 80 to 150 ml (\(p < 0.001\)).

The presence of LUTS, which were observed to be the most common postoperative low-grade AE (Clavien-Dindo grade I) decreased from 25% (\(n = 37\)) in 2011 to 8.8% (\(n = 7\)) in 2019 (\(p < 0.001\)) as shown in Fig. 3. Furthermore, postoperative hematuria (14.2–5%, \(p = 0.001\)) and urinary retention...
(6.1–0%, \( p = 0.035 \)) significantly decreased. All three of these postoperative AE remained constant across PV1-3. All other documented complications such as stroke, cardiac ischemia or angina pectoris stayed constant at an extremely low level with a total of 12 cases during the period of 2011–2019.

**Discussion**

The present GGG database analysis sought to evaluate the intraoperative progress in photoselective vaporization of the prostate with the GreenLight-XPS 180-Watt system (GL-XPS) in the largest documented cohort (> 3400) treated at...
seven global high-volume tertiary referral centers. Observing 3441 patients operated by eight surgeons with similar experience in performing this procedure at the start of our study strengthens the observations of intraoperative progress. We demonstrated a temporal improvement in the effectiveness, efficacy, and safety of GL-XPS.

One important aspect of progress with GL-XPS was the significant reduction in intraoperative and postoperative high-grade AEs while operative time and laser time showed a statistically significant increase but stayed highly efficient over the years of surgery and within the prostate PV groups.

First, our OT (> 55 min) and LT (> 27 min) between 2011 and 2019 is in line with data published by the multicenter GOLIATH study with an OT of 49.6 min (mean; SD 21.8) and a LT of 44.5 min (mean, SD 21.5) [6]. However, in comparison to the GOLIATH multicenter study, our LT was more efficient with a median time of 27 min (PV1, IQR 20–38) rising to a median of 75 min (PV3, IQR 54–96) for the larger glands sized close to 330 ml [10].

More interestingly the LT/OT ratio, as a parameter of intraoperative efficiency stayed significantly high with a value > 50% within each PV group from the beginning of this study and significantly improved during 2011–2019 to > 60%, independent to PV (PV1-3; p < 0.001). Our results are in line with multiple previously published results who postulated before, that larger PV prolongs the OT and LT a few minutes but not exponentially [11, 12]. Therefore, it was important to guarantee a high level of energy usage > 200 kJ and an energy density of 4 kJ/ml [10, 13].

Our database showed that the most important prognostic influencing factor for the significantly increasing parameters such as OT, LT, LT/ml or energy density is the PV. Furthermore, it is the most important factor for decreasing parameters such as preoperative and postoperative complications. Valdivieso et al. were able to show that GreenLight-XPS was feasible in men with very large prostates [13]. A more detailed analysis of the impact of the presence of a median lobe was recently published by the GGG showing that the presence of a median lobe has no clinically significant impact on procedural or postoperative outcomes for patients undergoing Greenlight PVP using the XPS-180 W system [14]. The increase in prostate volume and the significant increase in patients with a median lobe can be seen as a signal of the increasing surgical confidence in the Greenlight method for large prostate.

Progress in safety of GL-XPS is confirmed in our study with significant low intraoperative and postoperative AEs comparable with healthier patients on no permanent anticoagulation. Remaining on anticoagulation with for example acetylsalicylic acid or coumadin (35%, n = 1176, p = 0.907) while performing GL-XPS, had a low impact on morbidity or high-grade AEs [15].

Perioperative AEs significantly decreased over the years of surgery (p < 0.001) and showed in our sub-analysis a significant reduction in prostate glands up to 150 ml (p < 0.001) while complication-rate stayed constant on a low level in PV3. These results are in line with Nguyen et al.’s GGG database study which suggested GL-XPS to be safe in larger glands especially those with a median lobe [14] Likewise, Stone et al. and Meskawi et al. further confirmed GL-XPS’ safety for glands > 100 ml [4, 16]. Intraoperative transfusions being one of the most severe low-grade AE significantly (p = 0.004) dropped to 0% over time. This value is considered as a very low rate for glands > 100 ml [10].

The postoperative baseline results are in line with multicenter studies. A length of hospital stay with a median of

![Fig. 3 Early postoperative complications (< 30 days) between 2011 and 2019 in correlation to prostate volume group 1–3](image-url)
2 days, high symptom reduction in IPSS/QoL with median of 5/1, sufficient Qmax > 18 ml/s and a median catheterization time of 1 day stayed stable in our study between 2011 and 2019 and showed no difference within the different PV groups [17, 18].

Overall, postoperative AEs were significantly reduced to almost half across the time of our study. More specifically, the most common low-grade AE such as LUTS, hematuria, and urinary retention were reduced significantly between 2011 and 2019, while incontinence stayed constant over the observation period. These results are in line with Goueli et al. who showed significantly different postoperative rates in urinary retention vs. no retention collective comparison. Moreover, our results are confirmed as an extremely low AE rate even in the beginning of our study in 2011. The mod-est, yet steady reduction of AE rate between 2011 and 2019 further emphasizes the high level of surgeon experience at the start of the study with a potential in progress for each surgeon [19].

Overall, we assume with previous GGG results of Nguyen et al. that the surgeon’s level of experience with the GL-XPS 180-Watt laser system combined with a global experience transfer, together provides an efficient outcome for each high-volume center and a safe operation procedure for each patient [14].

Our GGG analysis is limited by its retrospective design, the short follow-up period of 12 months and that not all parameters were collected for all 3441 patients. Further, detailed parameters such as anticoagulants are incomplete in the GGG data base and therefore could not be analysed. Unfortunately, long term follow-up beyond 12 months is not available on a global scale yet. Each surgeon is already considered to be a highly experienced surgeon in GL-XPS and past their learning curve, operating for years, and may not be representative of an average or minimally experienced GL-XPS surgeon. Despite this limitation, our study is the first analysis to verify the perioperative progress of GL-XPS at a global scale.

Conclusion

Our GGG analysis showed that, within a 9-year time period, it is possible to significantly decrease intraoperative, transfusions and postoperative adverse events as part of the experience level progress of a surgeon across different prostate volume groups (especially up to 150 ml) while staying highly efficient in the operative time and laser time of GL-XPS.

Author contributions IL: data collection, data analysis, manuscript writing and editing. KL.: data collection, data analysis. KCZ: project development. HC: project development, manuscript editing. MR: manuscript writing and editing.

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Declarations

Conflict of interest Hannes Cash, Kevin Zorn, Dean Elterman, Ed-gardo F. Becher, Giovanni Ferrari and Luca Cindolo report receiving honoraria as consultant/proctor for Boston Scientific. All other authors report no conflict of interest.

Ethical approval All procedures performed in this study were in approval with the ethical standards of the institutional research commit-tee and with the 1964 Helsinki declaration and its later amendments or comparable ethical standards.

Consent for publication For this type of study, informed consent is not required (retrospective study).

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