

| Study, country | Study type, study period | Number of patients | Patient characteristics | Intervention | Comparison | Length of follow-up | Outcome measures and effect size | Additional comments | | | | | | | | | | | | | | | | | | | | |
|-------------------------------|--------------------------------------------------------------|---------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------|-------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------|----------------------------------------------------------------------------------------------------------------------------------------------------------------|--------------------------------------------------------------|---------------------|--------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------|-----------------------------|---------|-----------|--------|-----------|--------|-------------|----------|---------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------|----------|-----------------------------|--------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------|--------------|-----|------------|-------|-------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------|-----|------------------------------|-----------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------|---------------------------------|
| Srinivasan (1994) UK | Observational study (appears retrospective) 1982-1989 | 41 patients T3-4, Grade 2-3 TCC treated by palliative radiotherapy, presenting with haematuria and local pain | 19 patients with reasonable PS (WHO grade ≤3) treated with conventional palliative treatment; 22 patients with poor performance status (WHO grade ≥4) accelerated radiotherapy. Mean age 78.4 years in 2-fraction group compared to 71.6 yrs in conventional group. | Conventional palliative treatment 4500cGy in 12 fractions over 26 days Both regimens used supervoltage photons. From 1984 volume was localised with CT. | Accelerated radiotherapy 1700cGy in 2 fractions over 3 days. | Not reported | Improvement of pain: 73% (16/22) 2-fraction, 37% (7/19) conventional RT. Disease was fatal in all patients Overall survival: Mean 9.77 months 2-fraction vs 14.47 months conventional | No pain data for 7 patients | | | | | | | | | | | | | | | | | | | | |
| Kouloulias 2013 Greece | Prospective observational study 2005-2011 | 58 patients with organ-confined (cT1-2, N0) bladder cancer. All inoperable, with poor PS, >75yrs. Excluded previous pelvic RT or cystectomy, LN mets, distant mets or hip prosthesis. | <table border="1"> <tr> <td>Median age</td> <td>77 (70-91)</td> </tr> <tr> <td>T1</td> <td>12</td> </tr> <tr> <td>T2</td> <td>46</td> </tr> <tr> <td>PS 60-70%</td> <td>10</td> </tr> <tr> <td>PS 50-60%</td> <td>48</td> </tr> <tr> <td>Male/female</td> <td>47/11</td> </tr> </table> | Median age | 77 (70-91) | T1 | 12 | T2 | 46 | PS 60-70% | 10 | PS 50-60% | 48 | Male/female | 47/11 | Hypofractionated 3DCRT-virtual CT planning used. Clinical target volume (the bladder) and planning target volume obtained by expanding CTV with a margin of 1cm in each direction and of 0.5cm posteriorly. Entire bladder was treated using 4-field technique with 15 MV x-ray energy beams. 36Gy in 6 weekly fractions. | N/a | 3 months after RT treatment | Acute Grade 1-2 GI toxicity: 13/58 (22%) Acute Grade 1-2 GU toxicity: 19/58 (33%) No grade 3 or higher GI or GU toxicity. Patient-reported pain: VAS score improved from 4.2 (±1.1) to 1.8 (±0.6) (p<0.001). | | | | | | | | | |
| Median age | 77 (70-91) | | | | | | | | | | | | | | | | | | | | | | | | | | | |
| T1 | 12 | | | | | | | | | | | | | | | | | | | | | | | | | | | |
| T2 | 46 | | | | | | | | | | | | | | | | | | | | | | | | | | | |
| PS 60-70% | 10 | | | | | | | | | | | | | | | | | | | | | | | | | | | |
| PS 50-60% | 48 | | | | | | | | | | | | | | | | | | | | | | | | | | | |
| Male/female | 47/11 | | | | | | | | | | | | | | | | | | | | | | | | | | | |
| Caravatta 2012 Italy | Prospective observational study | 27 patients with locally advanced cancer and metastatic disease, ECOG PS ≤3, no previous RT to same region. | <table border="1"> <tr> <td>Median age</td> <td>72 (47-86)</td> </tr> <tr> <td>Male</td> <td>11 (41)</td> </tr> <tr> <td>Female</td> <td>16 (59)</td> </tr> <tr> <td>ECOG PS 0</td> <td>7 (26)</td> </tr> <tr> <td>PS 1</td> <td>6 (22)</td> </tr> <tr> <td>PS 2</td> <td>9 (33.5)</td> </tr> <tr> <td>PS 3</td> <td>5 (18.5)</td> </tr> <tr> <td>Primary cancer site</td> <td></td> </tr> <tr> <td>Gynaecologic</td> <td>48%</td> </tr> <tr> <td>Colorectal</td> <td>18.5%</td> </tr> </table> | Median age | 72 (47-86) | Male | 11 (41) | Female | 16 (59) | ECOG PS 0 | 7 (26) | PS 1 | 6 (22) | PS 2 | 9 (33.5) | PS 3 | 5 (18.5) | Primary cancer site | | Gynaecologic | 48% | Colorectal | 18.5% | Short course accelerated 3D conformal RT. CT planning used. Clinical target volume defined as primary tumour or metastatic site plus 1 cm margin. 1 cm margin in all directions added for planning target volume. | n/a | Median 6 months (range 3-28) | Acute Grade 1-2 GI toxicity: 5/27 Acute Grade 1-2 GU toxicity: 11/27 Patient-reported pain: 12 patients treated for pain control. Mean VAS score improved from 6 (±2) to 3 (±2.3) p=0.0002. 5 patients | Not all bladder cancer patients |
| Median age | 72 (47-86) | | | | | | | | | | | | | | | | | | | | | | | | | | | |
| Male | 11 (41) | | | | | | | | | | | | | | | | | | | | | | | | | | | |
| Female | 16 (59) | | | | | | | | | | | | | | | | | | | | | | | | | | | |
| ECOG PS 0 | 7 (26) | | | | | | | | | | | | | | | | | | | | | | | | | | | |
| PS 1 | 6 (22) | | | | | | | | | | | | | | | | | | | | | | | | | | | |
| PS 2 | 9 (33.5) | | | | | | | | | | | | | | | | | | | | | | | | | | | |
| PS 3 | 5 (18.5) | | | | | | | | | | | | | | | | | | | | | | | | | | | |
| Primary cancer site | | | | | | | | | | | | | | | | | | | | | | | | | | | | |
| Gynaecologic | 48% | | | | | | | | | | | | | | | | | | | | | | | | | | | |
| Colorectal | 18.5% | | | | | | | | | | | | | | | | | | | | | | | | | | | |

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| | | | Genitourinary | 33.5% | | | | | |
| | | 12 patients received RT for pain control. | | | Patients received 14Gy (3.5Gy fractions), 16Gy (4-Gy fractions), or 18 Gy (4.5Gy fractions) in 3 dose levels. Patients underwent RT on 2 consecutive days with twice-daily fractionation, with an interval of ≥8 hrs between fractions. | | | (41.7%) had complete pain relief, 6 patients (50%) showed more than 30% VAS reduction. 8 (66%) reported a reduction in pain score and 9 (75%) reported a reduced drug score. 1 of 4 patients discontinued opioid analgesic therapy. Quality of life (Cancer Linear Analog Scales): well-being, fatigue, ability to perform daily activities. 13 patients completed QOL VAS ranking. No significant differences between baseline and post-treatment, though improvement in all indexes was noted. | |
| Albers 2002 Germany | Prospective non-comparative phase 2 study 1998-1999 | N=30, proven, measurable recurrent or progressing TCC, prior cisplatin-based chemo. No prior gemcitabine, chemo or RT within 4 weeks prior to study, no karnofsky PS <40, adequate liver and renal function. | 86% prior radical surgery with adjuvant MVAC/MVEC or CM, 7% cystectomy with neoadjuvant MEC, 7% primary MVEC/MVAC without radical surgery 40% regional lymph node and distant metastases, 26% regional lymph nodes only 28 patients evaluable for response and toxicity, | | Gemcitabine 1250mg/m2 on day 1 & 8 of a 21-day course i.v. 30 mins. Maximum 6 courses (18 weeks) 9/28 completed 6 courses of treatment, 15 did not receive maximum number due to progression, 4 dropped out due to toxicity or personal reasons | n/a | Mean 8.4 months (0-25.3) | Toxicity: 36% (10/28) Grade 3-4 Leukocytopenia, 11% (3/28) thrombocytopenia, 11% (3/28) anemia, 11% (3/28) Grade 3 vomiting, 11% pulmonary toxicity, 3% (1/28) exanthema. 1 Grade 4 vomiting. Quality of life (Spitzer index values 10-point scale): In non-responders decreased from 7.8 ±2.4 to 6.7±2.2 at the end of treatment. In responders there was no change during treatment 8.0 ±1.6 before, 8.1 ±2.5 after. Pain scale (7-point scale): Non-responders showed decrease in pain values 5.3±1.8 to 4.8±1.5 (an increase in pain). Responders showed | QoL Spitzer questionnaire designed and validated for palliative care populations. |

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| | | | | | | | improvement in pain from 4.3±1.9 to 5.8 ±1.3 (p<0.05) | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | |
| Miyata 2012 Japan | Retrospective observational study 2003-2011 | 35 patients with metastatic and/or recurrent UC previously treated with cisplatin containing chemo | <table border="1"> <tr><td>Median age</td><td>68</td></tr> <tr><td>Male</td><td>26 (74%)</td></tr> <tr><td>Female</td><td>9 (25.7%)</td></tr> <tr><td>PS0</td><td>14 (40)</td></tr> <tr><td>PS1</td><td>16 (46)</td></tr> <tr><td>PS2</td><td>5 (14)</td></tr> <tr><td>Primary tumour site</td><td></td></tr> <tr><td>UUT</td><td>13 (37)</td></tr> <tr><td>Bladder</td><td>21 (60)</td></tr> <tr><td>both</td><td>1 (2.9)</td></tr> <tr><td>Prior treatment</td><td></td></tr> <tr><td>Chemo</td><td>8 (23)</td></tr> <tr><td>Chem+surgery</td><td>17 (49)</td></tr> <tr><td>Chem+Rad</td><td>6 (17)</td></tr> <tr><td>ChemRad+surgery</td><td>4 (11)</td></tr> <tr><td>2nd line CT</td><td>31 (89)</td></tr> </table> | Median age | 68 | Male | 26 (74%) | Female | 9 (25.7%) | PS0 | 14 (40) | PS1 | 16 (46) | PS2 | 5 (14) | Primary tumour site | | UUT | 13 (37) | Bladder | 21 (60) | both | 1 (2.9) | Prior treatment | | Chemo | 8 (23) | Chem+surgery | 17 (49) | Chem+Rad | 6 (17) | ChemRad+surgery | 4 (11) | 2 nd line CT | 31 (89) | All patients progressed after cisplatin based therapy. Low dose GP: Gem 700mg/m ² i.v. for 30mins on day 1 and 8 of each 28 day cycle. Pac at 70mg/m ² i.v. over 3h on day 1 and 8 of each 28-day cycle. Dexamthasone sodium phosphate (6.6mg), diphenhydramine hydrochloride (50mg) and ranitidine hydrochloride (100mg) were administered before treatment. Median 5 treatment cycles per patient. | n/a | Median 10 months (IQR 4-19) | Pain relief (VAS scale): Median abdominal/back pain median VAS score 4 (3-6) and all needed analgesics. After CT scores were 2 (1-3) (p<0.001). Improved pain scores (n=24, 69%), decrease in analgesic consumption (n=12, 34%). Decrease in analgesic consumption or decrease in VAS score without increasing analgesic dose 28/35 (80%). Toxicity: Grade 3-4 anemia (n=2, 6%), leukopenia (n=5, 14%), thrombocytopenia (n=2, 6%). Grade 3-4 Fatigue 0, nausea/vomiting (n=1, 3%), neuropathy 0, skin rash (n=1, 2.9%). | |
| Median age | 68 | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | |
| Male | 26 (74%) | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | |
| Female | 9 (25.7%) | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | |
| PS0 | 14 (40) | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | |
| PS1 | 16 (46) | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | |
| PS2 | 5 (14) | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | |
| Primary tumour site | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | |
| UUT | 13 (37) | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | |
| Bladder | 21 (60) | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | |
| both | 1 (2.9) | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | |
| Prior treatment | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | |
| Chemo | 8 (23) | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | |
| Chem+surgery | 17 (49) | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | |
| Chem+Rad | 6 (17) | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | |
| ChemRad+surgery | 4 (11) | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | |
| 2 nd line CT | 31 (89) | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | |
| De Leon-Casasola 1993 USA | Observational study (appears prospective) | 26 patients with pelvic pain from colorectal or gynaecological cancers that was no longer controlled with opioids or excessive sedation or side-effects from opioids. Excluded allergies to phenol, life expectancy <1 | <table border="1"> <tr><td>Mean age</td><td>55±8 years</td></tr> <tr><td>Gynae cancer</td><td>22 (77%)</td></tr> <tr><td>Prostate cancer</td><td>4 (15%)</td></tr> <tr><td>Colorectal</td><td>2 (7%)</td></tr> </table> | Mean age | 55±8 years | Gynae cancer | 22 (77%) | Prostate cancer | 4 (15%) | Colorectal | 2 (7%) | Hypogastric plexus block: Contrast medium used to determine accurate placement of needles. All underwent diagnostic block with 8ml 0.25% BUP injected through each needle. If 70% reduction in pain intensity a neurolytic block performed on following day. For neurolysis 8ml of 10% phenol (dissolved in sterile water) was used on each side. Criteria for success of | n/a | NR | Pain relief (Visual Analog Pain Scale 0-10): All patients 10/10 (worst pain) before block despite oral opioid therapy. Morphine sulphate mean was 953±722 mg/day before block to 420±354 2wks after block (p<0.0001). Patients in the success group were using sig less daily oral MS than patients in failure group (736±633 versus 1443 ±703 mg/day, p=0.02). % reduction in usage = 67% in success group and 45% in failure group. Overall 18 (69%) had satisfactory pain relief after 1 | No bladder cancer patients. Length of follow-up not reported. | | | | | | | | | | | | | | | | | | | | | | | | |
| Mean age | 55±8 years | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | |
| Gynae cancer | 22 (77%) | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | |
| Prostate cancer | 4 (15%) | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | |
| Colorectal | 2 (7%) | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | |

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| | | mo, patients receiving RT/CT | | block: 1)decrease in VAPS of at least 70% or pain intensity of less than 3 to 10 during first two weeks after neurolytic block. 2)decrease in opioid requirements of at least 30% resulting in disappearance of bothersome side-effects 2 wks after block. Patients with failed blocks after 2 consecutive attempts received continuous epidural BUP morphine. | | | or 2 procedures with a VAPS of ≤ 3 . Complications; No intraoperative complications. No long-term complications. | | | | | | | | | | | | | | | | | | | | | |
| Gamal 2006 Egypt | Prospective observational study | 30 patients with pelvic pain due to cancer and had been treated with analgesic medication according to WHO guidelines and still had pain on VAS >4 (0-10, worst pain) | <table border="1"> <tr> <td></td> <td></td> </tr> <tr> <td>Male</td> <td>14 (47%)</td> </tr> <tr> <td>female</td> <td>16 (53%)</td> </tr> <tr> <td>Mean age</td> <td>59</td> </tr> <tr> <td>Pain duration (mo)</td> <td>5.4-6.1</td> </tr> <tr> <td colspan="2">Primary cancer site</td> </tr> <tr> <td>Rectum</td> <td>5 (16%)</td> </tr> <tr> <td>Cervix</td> <td>8 (27%)</td> </tr> <tr> <td>Bladder</td> <td>9 (30%)</td> </tr> <tr> <td>endometrial</td> <td>8 (27%)</td> </tr> </table> <p>Patients with no contraindications to regional blockade and sympathetic blockade.</p> | | | Male | 14 (47%) | female | 16 (53%) | Mean age | 59 | Pain duration (mo) | 5.4-6.1 | Primary cancer site | | Rectum | 5 (16%) | Cervix | 8 (27%) | Bladder | 9 (30%) | endometrial | 8 (27%) | Superior hypogastric block: patients randomised into two groups - transdiscal approach (n=15) or block via classic posterior approach (n=15). | Transdiscal versus posterior approach | 3 months | VAS pain scores: Scores decreased from baseline at 24h, 1 week, 1 month and 2 months after block ($p < 0.05$). At 3 months there was no difference from baseline. No differences between the two groups at any timepoint. Daily morphine consumption decreased in both groups from baseline up to 2 months after block. No differences between 3 month and baseline. Complications: In the classic group: Intravascular puncture (n=2, 13%), urinary injury (n=4, 27%) | No details about randomisation to two groups. |
| | | | | | | | | | | | | | | | | | | | | | | | | | | | | |
| Male | 14 (47%) | | | | | | | | | | | | | | | | | | | | | | | | | | | |
| female | 16 (53%) | | | | | | | | | | | | | | | | | | | | | | | | | | | |
| Mean age | 59 | | | | | | | | | | | | | | | | | | | | | | | | | | | |
| Pain duration (mo) | 5.4-6.1 | | | | | | | | | | | | | | | | | | | | | | | | | | | |
| Primary cancer site | | | | | | | | | | | | | | | | | | | | | | | | | | | | |
| Rectum | 5 (16%) | | | | | | | | | | | | | | | | | | | | | | | | | | | |
| Cervix | 8 (27%) | | | | | | | | | | | | | | | | | | | | | | | | | | | |
| Bladder | 9 (30%) | | | | | | | | | | | | | | | | | | | | | | | | | | | |
| endometrial | 8 (27%) | | | | | | | | | | | | | | | | | | | | | | | | | | | |
| Plancarte 1997 Mexico/USA | Prospective observational study | 227 patients with pelvis pain and gynaecologica l, colorectal or | Excluded patients with anticoagulopathies, allergies to phenol, life expectancy <3 months, concurrent rad/CT or scheduled to receive treatment within 4 wks of | Superior hypogastric plexus block: L4-L5 intervertebral space was found and marked. Patient sedated. Accurate placement of | n/a | 6 months | Patient-reported pain: Preblock VAS score 8-10 (n=159, 100%). No patients had score 8-10 post-block. Postblock VAS score 4-7 (n=60, | Oral morphine therapy not available in Mexico when | | | | | | | | | | | | | | | | | | | | |

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| | | genitourinary cancer who had poor pain control – failed opioid therapy or excessive side-effects/sedation. | <p>block, purely somatic and/or neuropathic pain.</p> <p>159 (79%) had a positive response to diagnostic block and therefore available for neurolytic block.</p> <p>Mean age =48 years. 89% were women with gynaecological cancers, 17 males with prostate, colorectal or bladder carcinoma.</p> | <p>needles determined by collection of contrast medium just anterior to the L5-S1 intervertebral space. All patients had diagnostic block with 8ml of 0.25% bupivacaine injected through each needle. If patient reported a 50% reduction in pain intensity for 4 hrs, a neurolytic block was done on the following day. If patients failed to derive a benefit from this technique they were removed from study and offered epidural analgesia. For neurolysis 8ml of 10% phenol used on each side, dissolved in sterile water.</p> | | | <p>38%), score <4 (n=99, 62%). Overall 115/159 (72%) who responded to diagnostic block had satisfactory pain relief after one or two procedures. This is 51% (115/227) overall response for eligible patients</p> <p>Complications: No intraoperative complications, no long-term complications.</p> | <p>study was done.</p> <p>Mostly gynaecological cancers.</p> |
| Plancarte 1990 USA/Mexico | Prospective observational study | 28 patients – 22 female , 5 male | <p>Mean age 36 years. All had chronic lower abdominal pain with a prominent visceral component, secondary to advanced cancer. (20 cervix, 4 prostate, 1 testicle, 1 post-radiation cystitis) Persistent pain despite radiotherapy, chemotherapy, non-opioid and opioid analgesics, and behavioural pain management.</p> | <p>Superior hypogastric plexus block: Location of L4-L5 interspace was located. Block used for either diagnostic/prognostic or therapeutic purposes. For diagnosis 6-8ml 0.25% bupivacaine was used. For therapeutic (neurolytic) blocks a total of 6-8ml 50% alcohol through each needle</p> | n/a | Monthly until death | <p>Visual and verbal analogue scales used to measure pain immediately before block and after at 30 mins, 1,2,4,5 and 24 hr, then monthly until death.</p> <p>Patient-reported pain: Mean reduction in pain of 70% was observed, residual pain seemed generally of somatic origin. Injections of epidural steroids, serial injections of 2-3% epidural phenol and or non-opioid analgesics used to control remaining somatic pain resulting in a global reduction of pain scores by 90%. No</p> | <p>Outcomes poorly reported. No mean scores. No bladder cancer patients.</p> |

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| | | | | | | | return of pain in all but two patients. | |
| Cariati 2002 Italy | Observational study (appears prospective) 1995-2000 | 10 patients with pelvic malignancy and chronic severe pain which could be controlled only with high doses of NSAIDs | 4 rectum/sigmoid adenocarcinoma, 3 uterine carcinoma, 2 bladder cancer, 1 secondary lesion of the right hip from laryngeal carcinoma. Nine treated with morphine sulphate, one with high levels of NSAIDs | CT-guided superior hypogastric neurolytic block with alcohol, using a single needle and anterior approach. All patients sedated before procedure. The first four patients had an injection of 10ml alcohol, 2 had 15ml and the last 4 patients had 20ml alcohol. | n/a | Not reported | Complications: No local complications such as hematomas, or puncture of vascular stricture. Pain: evaluated using a 4-grade subjective analogue scale (none, mild, moderate, complete). 4/10 Complete disappearance of pain lasting until patient death (60-160 days after procedure) and with no analgesics, 2/10 moderate reduction with no opioids, 3/10 mild reduction of pain (one died at 20 days, 2 had opioids restarted at 17 and 25 days). One patient had no benefit and restarted opioid treatment. | Mostly colorectal or uterine cancer. |