

Supplementary material 1. Characteristics of included randomized controlled trials

First author (year)	Participants and setting	Interventions	Outcomes
Apuzzio (1989) ¹⁸	<p>Number of women randomized: 25 for PID</p> <p>Number of women analyzed: group A: 10; group B: 15</p> <p>Age (years): group A: 23.2; group B: 23.2</p> <p>Number of centres (country): 1 (US)</p>	<p>Group A: ciprofloxacin 300 mg IV every 12 h; treatment continued for 3-5 days until the woman was asymptomatic for 24 h. Upon discharge from hospital, women received oral antibiotics to complete 10-14 days of ciprofloxacin 750 mg PO twice daily</p> <p>Group B: clindamycin 900 mg IV every 8 h + gentamicin 1.5 mg/kg IV every 8 h. Upon discharge from the hospital, the women received oral antibiotics to complete 10-14 days of clindamycin 450 mg PO every 6 h</p>	<p>Primary outcome: treatment success, defined as women asymptomatic for 24h.</p>
Arredondo (1997) ²⁴	<p>Number of women randomized: 69 in each group</p> <p>Number of women analyzed: 69 in each group</p> <p>Age (mean) (years): group A: 28.9; group B: 30.7</p> <p>Number of centres (country): 6; Chile (1 centre), Peru (2 centres), Colombia (2 centres), and Mexico (1 centre)</p>	<p>Group A: clindamycin 300 mg (2 capsules 3 times daily) + ciprofloxacin (250 mg, 1 tablet twice daily) for 14 days and placebo IM (for an equivalent of 1 dose of ceftriaxone)</p> <p>Group B: ceftriaxone 250 mg IM (as a single dose) + doxycycline 100 mg (1 capsule twice daily) and placebo (2 capsules 3 times daily for equivalent doses of clindamycin) for 14 days</p>	<p>Primary outcome: clinical cure defined by the absence of, or minimal, pelvic tenderness, temperature <37.5 °C, and a WBC count of 10,000/mm³, if a minimum of 4 days of treatment had been completed. Clinical improvement defined as resolution of 2 of these 3 symptoms.</p> <p>Secondary outcomes: failure when 1 of the following circumstances was noted after at least 48 h of protocol therapy: signs and symptoms remained unchanged or worsened (during the first 72 h of therapy).</p> <p>Microbiological cure defined as eradication of <i>Neisseria gonorrhoeae</i> or <i>Chlamydia trachomatis</i> (or both) from clinically cured women. Failure defined as persistence of 1 or both of these 2 organisms or, in the case of clinical improvement or failure, the presence of endocervical pathogens. Superinfection defined as the isolation of ≥ 1 new pathogens.</p> <p>Adverse effects leading to discontinuation of treatment.</p>
Acioglu (2013) ¹⁹	<p>Number of women randomized: 1156</p> <p>Number of women analyzed: group A: 578; group B: 578</p>	<p>Group A: moxifloxacin 400 mg once daily for 14 days</p> <p>Group B: ofloxacin 400 mg twice daily + metronidazole 500 mg PO twice daily</p>	<p>Primary outcome: clinical cure, defined as a ≥ 60% reduction in the total pain score at day 21 compared with baseline and the absence of pelvic discomfort and tenderness,</p>

	Age (mean \pm SD) (years): group A: 30.3 \pm 3.7; group B: 29.3 \pm 3.5 Number of centres (country): 4 (Turkey)		temperature < 37.8 °C, and WBC < 10,000/mm ³ on day 21. Secondary outcomes: microbiological cure, adverse effects.
Balbi (1996) ⁵³	Number of women randomized: 78 Number of women analyzed: 76; group A: 40; group B: 36 Age (mean \pm SD) (years): group A: 25.3 \pm 7.7.4; group B: 29.4 \pm 7.8 Number of centres (country): 1 (Italy)	Group A: gentamicin, 2 mg/kg IV (attack dose), followed by 1.5 mg/kg IV every 8 h + clindamycin 900 mg IV every 8 h for 4 days, followed by clindamycin 450 mg PO every 6 h for a total of 14 days of treatment Group B: ceftazidime 1 g IV every 8 h + doxycycline 100 mg PO every 12 h for 4 days, followed by doxycycline 100 mg PO every 12 h for a total of 14 days of treatment	Primary outcome: clinical recovery, defined as: body temperature < 37 °C per 48 h, disappearance of pelvic pain, no increase of eventual adnexal mass after 7 days of the end of treatment Secondary outcome: follow-up performed 30 days after treatment finished; endocervical culture for <i>Neisseria gonorrhoeae</i> and <i>Chlamydia trachomatis</i> and endometrial culture for <i>Chlamydia trachomatis</i> performed in all positive cases at admission.
Bevan (2003) ³¹	Number of women randomized: 310 Number of women analyzed: 309; group A: 106; group B: 107; group C: 96 Age (mean (range)) (years): group A: 28.4 (18-54); group B: 27.6 (18-46); group C: 27.6 (17-54) Number of centres (country): multiple, but the authors did not specify how many (UK and others that were not stated)	Group A: azithromycin 500 mg IV single dose, days 2-7: 250 mg PO once daily or days 1-2: azithromycin 500 mg IV once daily; days 3-7: 250 mg PO once daily Group B: as group A + day 1: metronidazole 500 mg IV 3 times daily or 400 mg PO 3 times daily, days 2-12: metronidazole 400mg PO 3 times daily or days 1-2: azithromycin 500 mg IV once daily; days 3-7: 250 mg PO once daily plus day 1-2: metronidazole 500 mg IV 3 times daily, or 500 mg PO 3 times daily. Days 3-12: metronidazole 500 mg PO 3 times daily or days 1-21: metronidazole 500 mg PO 3 times daily Group C: day 1: metronidazole 500 mg IV 3 times daily, or metronidazole 400 mg PO 3 times daily; day 2-12: metronidazole 400 mg PO 3 times daily + days 1-14: cefoxitin 2 g IV or IM 4 times daily + day 1: probenecid 1 g PO single dose or day 1-21: doxycycline 100 mg PO twice daily + day 1-5: amoxicillin/clavulanate 1 g IV + times daily; day 6-21 amoxicillin/clavulanate 500 mg PO 3 times daily	Primary outcome: clinical response to treatment: cure, resolution of all baseline signs and symptoms; improvement, lessening of the baseline signs and symptoms or absence of \geq 1, but not all, of the baseline findings; or failure, no improvement or deterioration of baseline condition. Successful clinical outcome defined as cure or improvement. Assessment on day 15 (9-26 inclusive) and at follow-up (day 35-44). Microbiological outcome: eradication, absence of the baseline isolate(s); persistence, presence of baseline isolate(s); or superinfection, presence of a micro-organism different from that found at baseline.
Buisson (1989) ³⁸	Number of women randomized: 82 Number of women analyzed: group A: 42; group B: 40	Group A: amoxicillin-clavulanic acid 1 g IV every 8 h for at least 48 h, then 1.5-2 g PO twice daily. Mean length of treatment 19 days, never less than 14 days	Primary outcomes: clinical cure; defined as absence of fever, pain, and previously observed adnexal masses at 5-8 weeks'

	Age (mean (range)) (years): group A: 27.7 (18-46); group B: 28.7 (15-49) Number of centres (country): 8 (France)	Group B: amoxicillin 3-4 g IV per 24 h, mean 4 days, then 1.5-2 g PO daily. Mean length of treatment 17 days + aminoside (chosen by researcher's preference) 3-5 mg/kg IM per 24 h 2 or 3 times daily depending of the aminoside, mean length of treatment 7 days + metronidazole 1 or 1.5 g IV or suppository daily. For each case, a secondary prescription for a tetracycline 200 mg per 24 h was given, either immediately if results of laparoscopy or other investigations justified it, or later on positive <i>Chlamydia trachomatis</i> serology. Length of this treatment 3-4 weeks as decided by the researcher	follow-up; adverse events leading to discontinuation of treatment
Burchell (1987) ²⁸	Number of women randomized: 30 women, 10 in each group Number of women analyzed: 30 women, 10 in each group Age (years): not stated. Number of centres (country): not stated (South Africa)	Group A: doxycycline infusion 200 mg in 200 ml 5% dextrose over 2 h + doxycycline 100 mg after 24 h. The course was completed with oxytetracycline 250 mg every 6 h for 14 days Group B: ampicillin + metronidazole every 6 h 1 g IV and then 500 mg PO every 6 h for 14 days Group C: tetracycline + metronidazole with 3 × 1 g suppositories every 8 h and then 400 mg PO every 8 h for 14 days	Primary outcome: clinical cure
Ciraru-Vigneron (1986) ³⁴	Number of women randomized: 44; 22 in each group Number of women analyzed: 44; 22 in each group Number of centres (country): 1 (France) Age (years): not stated	Group A: amoxicillin-clavulanic acid while in hospital, 4 g per 24 h, first by IV, then PO once symptoms improved. At discharge, amoxicillin-clavulanic acid PO if <i>Chlamydia trachomatis</i> serology was negative; if positive for <i>Chlamydia trachomatis</i> , doxycycline prescribed for 3 weeks (dose not stated) Group B: ampicillin 6 g per 24 h IV+ gentamicin 160mg per 24 h IM+metronidazole 1.5 g per 24 h IV; when switched to oral administration, ampicillin replaced by amoxicillin 3 g per 24 h + metronidazole 1.5 g per 24 h PO and gentamicin 160 mg per 24 h IM for minimum of 7 days. At discharge, amoxicillin-clavulanic acid PO if <i>Chlamydia trachomatis</i> serology negative; if positive for <i>Chlamydia trachomatis</i> , doxycycline for 3 weeks (dose not stated)	Primary outcomes: clinical cure, defined by no fever, reduction of pain, and normal WBC count at discharge and in 30 days; adverse events leading to discontinuation of treatment Secondary outcomes: microbial cure of <i>Neisseria gonorrhoeae</i> and <i>Chlamydia trachomatis</i> ; length of hospital stay
Ciraru-Vigneron (1989) ³⁵	Number of women randomized: 165 women; group A: 78; group B: 87	Group A: amoxicillin 1 g IV + clavulanic acid 200 mg (Augmentin 1.2 g), 3 or 4 times daily until clinical and laboratory findings	Primary outcomes: excellent response defined as resolution of physical findings with continued improvements in laboratory values;

	<p>Number of women analyzed: 152 women; group A: 70; group B: 82 Number of centres (country): 8 (France) Age (mean) (years): group A: 26.9; group B: 25.1</p>	<p>improved, after which 4-6 tablets containing amoxicillin 500 mg + clavulanic acid 125 mg (Augmentin 625 mg) per tablet</p> <p>Group B: amoxicillin or ampicillin 3-4 g daily, with an aminoglycoside (gentamicin 160 mg, dibekacin 150 mg, or tobramycin 150 mg) + metronidazole 1.5 g daily parenterally. Subsequent conversion was to oral combination of amoxicillin or ampicillin 2-3 g + metronidazole 1-1.5 g daily</p>	<p>favourable response equate with a favourable course, allowing the persistence of ≥ 1 clinical signs or abnormal laboratory values (or both); failure defined as absence of therapeutic efficacy or of a favourable course after at least 6 days of therapy that a change in management, either surgery or an alternative antibiotic treatment was warranted</p>
Crombleholme (1986) ³⁶	<p>Number of women randomized: 39; group A: 20; group B: 19 Number of women analyzed: 39; group A: 20; group B: 19 Age (years): not stated. Number of centres (country): not stated (not stated)</p>	<p>Group A: sulbactam 1 g + ampicillin 2 g IV every 6 h</p> <p>Group B: metronidazole 15 mg/kg loading followed by 7.5 mg/kg IV every 6 h and gentamicin 1.5 mg/kg IV every 8 h</p> <p>Antibiotics in both groups continued for minimum of 5 days</p>	<p>No relevant outcomes reported for our analysis because the outcomes were not separately reported for the different diagnoses within the study groups. Authors stated clinical cure occurred in 19/20 women in group A and 16/19 women in group B</p>
Crombleholme (1987) ³⁷	<p>Number of women randomized: 44; 22 in each group Number of women analyzed: 42; 21 in each group Age (mean \pm SD) (years): group A: 27.7 \pm 6.9; group B: 29.0 \pm 12.1 Number of centres (country): 1 (US)</p>	<p>Group A: ampicillin 2 g + sulbactam 1 g IV every 6 h.</p> <p>Group B: metronidazole 15 mg/kg IV every 6 h + gentamicin 1.5 mg/kg IV every 8 h</p> <p>Therapy continued until women became afebrile and were without clinical signs of infection for 48 h or until clinical judgement dictated cessation of therapy. Range of treatment 3-11 days.</p>	<p>Primary outcome: clinical cure defined as absence of fever, without clinical signs of infections for 48 h or until clinical judgement</p>
Crombleholme (1989) ⁴⁴	<p>Number of women randomized: 80; 40 in each group Number of women analyzed: 80; 40 in each group Age (mean \pm SD) (years): group A: 25.7 \pm 4.8; group B: 25.3 \pm 6.2 Number of centres (country): 1 (US)</p>	<p>Group A: ciprofloxacin 300 mg IV every 12 h for 2-5 days, with ≥ 2 days of parenteral therapy and 48 h without fever before switching to ciprofloxacin 750 mg PO every 12 h, to complete a 14-day course</p> <p>Group B: clindamycin 600 mg IV every 6 h + 1 mg/kg IV every 8 h after ≥ 4 days of parenteral gentamicin and 48 h without fever before switching to clindamycin 300 mg PO every 6 h, to complete a 14-day course</p>	<p>Primary outcome: clinical cure at 3 days: improvement in clinical signs and symptoms Secondary outcomes: microbial cure of <i>Chlamydia trachomatis</i> and <i>Neisseria gonorrhoeae</i></p>
European Study Group (1992) ⁵⁰	<p>Number of women randomized: 170 women; group A: 88; group B: 82 Number of women analyzed: 170 women; group A: 88; group B: 82</p>	<p>Group A: clindamycin 900 mg IV every 8 h + gentamicin 2 mg/kg IV, followed by 1.5 mg/kg IV every 8 h for minimum of 4 days. At the end of the period of IV therapy, clindamycin 450 mg PO every 6 h was given to complete 14 days of treatment</p>	<p>Primary outcomes: clinical failure, minimum of 48 h of protocol therapy and characterized by signs and symptoms as unchanged or worsened during the first 48-72 h of</p>

	Age (mean) (years): whole study group: 28; age per group not stated. Number of centres (country): 10 (9 in Europe and 1 Africa)	Group B: cefoxitin 2 g IV every 6 h + doxycycline 100 mg IV every 12 h were given for at least 4 days. At the end of IV therapy, doxycycline 100 mg PO was given every 12 h to complete a total of 14 days of treatment	treatment, or worsening later, failure to improve further; need of additional antibiotics or need for surgery considered as failure; adverse events leading to discontinuation of therapy Secondary outcomes: microbial cure of <i>Chlamydia trachomatis</i> and <i>Neisseria gonorrhoeae</i>
Fischbach (1994) ³³	Number of women randomized: 60 Number of women analyzed: 57; group A: 26; group B: 31 Age (average (range)) (years): group A: 27 (30 ± 10); group B: 22 (26 ± 7) Number of centres (country): not stated (Germany)	Group A: ciprofloxacin 2 × 0.2 g IV daily + metronidazole 3 × 0.5 g IV daily Group B: cefoxitin 3 × 2 g IV daily + doxycycline 2 × 0.1 g IV daily After 2-5 days of treatment, ciprofloxacin, metronidazole, and doxycycline given PO. Both groups treated for 7-14 days	Primary outcome: clinical cure defined as subjective lack of symptoms, improvement upon gynaecological examination, normal leukocytes count, declining ESR, no fever, and elimination of bacteria
Giraud (1989) ³⁹	Number of women randomized: 152 Number of women analyzed: group A: 70; group B: 82 Age (mean) (years): group A: 26.9; group B: 25.1 Number of centres (country): unclear (France)	Group A: amoxicillin-clavulanic acid; while in hospital, 3 or 4 g per 24 h, by IV or perfusion, then 2 or 3 g per 24 h PO Group B: parenteral: ampicillin 3 or 4 g IV per 24 h, OR amoxicillin 3 g IV per 24 h + gentamicin 160 mg IM per 24 h OR dibekacin 150 g IM per 24 h + metronidazole 1.5 g IV per 24 h. Oral administration: ampicillin 3 g per 24 h OR amoxicillin 2 g/24 h + metronidazole 1.5 or 2 g per 24 h Additional treatments were limited, but could include: surgical or laparoscopic drainage of pus, local antibiotic for treatment of trichomonacides, non-steroidal anti-inflammatory drugs, corticoids	Primary outcomes: clinical cure at 10th day; clinical progress and improvement of biological parameters Secondary outcome: length of hospital stay
Heinonen (1989) ⁴⁰	Number of women randomized: 40 Number of women analyzed: group A: 16; group B: 20 Age (mean ± SD (range)) (years): group A: 29 ± 8 (18-43); group B: 29 ± 1 (16-50) Number of centres (country): 1 (Finland)	Group A: ciprofloxacin 200 mg IV every 12 h for 2 days followed by 750 mg PO every 12 h to complete 14-day course Group B: doxycycline 100 mg IV every 12 h + metronidazole 500 mg IV every 8 h for the first 2 days, followed by doxycycline 150 mg PO every 24 h + metronidazole 400 mg PO every 8 h to complete a 14-day course	Primary outcomes: clinical response based on a scale from 0 (absent or normal) to 3+ (severe) assessed on days 3, 6, 14, and 21 after the antimicrobial treatment was started. Failure defined as the presence of ≥ 1 of following criteria: no improvement in the clinical severity score at day 3 after the microbial treatment was started; CRP > 20 mg/L or a decline < 50% in the initial CRP level

			at day 6; positive cervical culture of <i>Neisseria gonorrhoeae</i> or <i>Chlamydia trachomatis</i> at days 14 or 21; or the need for additional antimicrobial agents or surgical intervention Secondary outcomes: adverse reactions and effects
Hemsell (1994) ⁵²	Number of women randomized: 344 Number of women analyzed: group A: 109; group B: 110; group C: 108 Age (mean ± SD) (years): success group: 24.7 ± 4.9; failure group: 26.2 ± 5.7 Number of centres (country): 6 (US)	Group A: cefoxitin 2 g IV every 6 h + doxycycline 100 mg every 12 h; followed by doxycycline 100 mg PO twice daily for a total of 10-14 days Group B: clindamycin 900 mg IV + gentamicin 1.5 mg/kg every 8 h after an initial gentamicin loading dose calculated at 2 mg/kg followed by clindamycin 450 mg PO 4 times daily for a total of 10-14 days Group C: cefotetan 2 g IV + doxycycline 100 mg every 12 h, followed by doxycycline 100 mg PO twice daily for a total of 10-14 days	Primary outcome: clinical cure defined as reduction of the severity score by ≥ 70%, with a normal temperature and leukocyte count
Heystek (2009) ⁴¹	Number of women randomized: 686 Number of women analyzed: 669; group A: 343; group B: 326 Age (mean ± SD) (years): group A: 29.0 ± 7.3; group B: 28.2 ± 7.2 Number of centres (country): 43 (14 countries)	Group A: moxifloxacin 400 mg PO once daily for 14 days Group B: doxycycline 100 mg PO twice daily + metronidazole 400 mg PO 3 times daily for 14 days + ciprofloxacin 500 mg PO once daily for 14 days	Primary outcome (modified ITT): clinical cure at 2-14 days' post treatment: clinical success defined as cure (severity score reduced by ≥70% plus normal temperature and leukocyte count) or improvement (severity score reduced <70% but >30% plus normal temperature and leukocyte count). Therapy considered to have failed if symptoms and signs of infection persisted or worsened, as shown by persistent fever, leukocytosis, a reduction in severity score of ≤30%, or a combination of these. Clinical efficacy was 'unevaluable' when a woman could not be assessed Secondary outcome (PP): microbiological clearance of <i>Chlamydia trachomatis</i> , microbiological clearance of <i>Neisseria gonorrhoeae</i>

<p>Hoyme (1993) ⁵⁴</p>	<p>Number of women randomized: 33 Number of women analyzed: 33; group A: 15; group B: 18 Age (years): not stated Number of centres (country): 1 (Germany)</p>	<p>Group 1: ofloxacin 2 × 200 mg + metronidazole 2 × 500 mg, first IV and then PO for 10 days in total Group 2: gentamicin 3 × 80 mg + clindamycin 4 × 600 mg (initially 1200 mg IV) for 10 days in total</p>	<p>Primary outcome: clinical cure, no raw data reported for outcomes</p>
<p>Judlin (2010) ²⁵</p>	<p>Number of women randomized: 460 Number of women analyzed: ITT: group A: 228; group B: 232; PP: group A: 194; group B: 190 Age (mean ± SD) (years): group A: 35.2 ± 8.4; group B: 35.4 ± 8.7 Number of centres (country): 7 (China, Indonesia, South Korea, The Philippines, Pakistan, Thailand, Taiwan)</p>	<p>Group A: moxifloxacin 400 mg PO once daily for 14 days Group B: levofloxacin 500 mg (2 x 250 mg tablets) PO once daily + metronidazole 500 mg (1 tablet) PO twice daily for 14 days</p>	<p>Efficacy: primary efficacy variable was clinical response at test-of-cure in the PP population: 'Clinical success' defined as women with clinical cure at test-of-cure; failures were women with failure at the 'during therapy' visit or improvement or failure at test-of cure. 'Clinical cure' defined as reduction in tenderness score (McCormack scale) of >70%, apyrexia (rectal/tympanic/oral temperature < 38.0 °C or axillary temperature <37.5 °C) and WBC < 10,500/mm³. 'Clinical improvement' defined as reduction in tenderness score of 30-70%, apyrexia (rectal/tympanic/oral temperature < 38.0 °C or axillary temperature < 37.5°C) and WBC < 10,500/mm³. 'Clinical failure' defined as reduction in tenderness score of < 30% or elevated temperature (rectal/tympanic/oral temperature ≥ 38.0 °C or axillary temperature < 37.5 °C) or WBC ≥ 10,500/mm³, or a combination of these. Secondary efficacy variables: Clinical response during therapy (PP population) classified as 'clinical improvement', defined as reduction in tenderness score of > 30% with improvement in temperature, or 'clinical failure', defined as persistence or worsening of symptoms and signs of infection, as evidenced by a reduction in tenderness score of > 30% or no improvement in temperature (or both)</p>

Landers (1991) ⁵¹	<p>Number of women randomized: 162 Number of women analyzed: 148, group A: 75; group B: 73 Age (mean \pm SD) (years): total: 23.5 \pm 6.1; group A: 23.3 \pm 5.3; group B: 23.8 \pm 6.0 Number of centres (country): 2 (US)</p>	<p>Group A: cefoxitin 2 g IV every 6 h + doxycycline 100 mg IV every 12 h for minimum 4 days and at least 48 h after disappearance of fever. Women without fever on admission treated for minimum of 4 days in the hospital for at least 24 h beyond the adequate relief of pain and tenderness to a normal lifestyle without surgical intervention. After discharge from the hospital, doxycycline 100 mg PO twice daily was continued to complete a total of 14 days of treatment</p> <p>Group B: clindamycin 600 mg IV every 6 h + tobramycin 2 mg/kg IV for 1 dose, followed by 1.5 mg/kg IV every 8 h for minimum of 4 days and for at least 48 h after disappearance of fever. Women without fever on admission treated for minimum of 4 days in hospital for at least 24 h beyond adequate relief of pain and tenderness to a normal lifestyle without surgical intervention. After discharge from hospital, clindamycin 450 mg PO 4 times daily continued to complete a total of 14 days of treatment</p>	<p>Primary outcome (ITT): clinical response for a satisfactory initial clinical response defined as an improvement of admitting signs and symptoms, included abdominal-pelvic pain, fever, and pelvic tenderness. Follow-up evaluation performed at hospital discharge and at 2-6 weeks after initial enrolment Secondary outcomes (PP): microbiological clearance of <i>Chlamydia trachomatis</i> and reduction in tenderness score</p>
Leboeuf (1987) ⁴²	<p>Number of women randomized: 45; group A: 23; group B: 22 Number of women analyzed: 39; group A: 21; group B: 18 Age (mean \pm SD) (years): group A: 27.9 \pm 5.2; group B: 29.1 \pm 9.5 Number of centres (country): 2 (France)</p>	<p>Group A: clindamycin 900mg IV every 8 h diluted in 150mL (minimum volume) saline slow perfusion (30-60 minutes) + gentamicin 1 mg/kg IM every 8 h (with minimum dose prescribed of 60 mg every 8 h, according to bodyweight). Treatment given for minimum of 5 days in hospital. Clindamycin perfusion not stopped until 48 consecutive h with temperature below 37.5 °C; at that point treatment could be PO. Maximum length of treatment at the discretion of the therapist. If also treated with a tetracycline, this was not prescribed < 48 h after the end of the treatment protocol (either clindamycin + gentamicin or metronidazole + gentamicin)</p> <p>Group B: metronidazole 500 mg every 8 h in slow IV perfusion (30-60 minutes) + gentamicin 1 mg/kg IM every 8 h (with minimum dose prescribed of 60 mg every 8 h, according to bodyweight). Treatment given for 6 weeks</p>	<p>Primary outcome: clinical cure: absence of infection in the days following cessation of treatment according to clinical observations, microbe eradicated during or after treatment Secondary outcome: length of hospital stay</p>
Malhotra (2003) ²⁰	<p>Number of women randomized: 165 Number of women analyzed: 153 women; group A: 52; group B: 48; group C: 53</p>	<p>Group A: ciprofloxacin 500mg + tinidazole 600mg twice daily for 7 days</p> <p>Group B: fluconazole 150 mg (1 tablet) + azithromycin 1 g (1 tablet) + secnidazole 2 g (2 tablets). Advised to take azithromycin</p>	<p>Primary outcome: clinical cure defined as at least 70% reduction in severity score, no more than mild abdominal pain, and no recurrence of symptoms or signs of PID within 4 weeks of</p>

	Age (mean \pm SD) (years): group A: 25.6 \pm 3.9; group B: 25.8 \pm 3.2; group C: 25.5 \pm 4.9 Number of centres (country): 1 (India)	on empty stomach in the morning, secnidazole with or after food and fluconazole in the evening Group C: doxycycline 100 mg twice daily + metronidazole 200 mg 3 times daily for 1 week. The 2 drugs were available in the hospital pharmacy free of cost	therapy. Treatment failure defined as < 20% decrease in tenderness score
Martens (1990) ⁴⁹	Number of women randomized: 99 Number of women analyzed: 94 Age (mean \pm SD) (years): group A: 26 \pm 7; group B: 25 \pm 7; group C: 26 \pm 7 Number of centres (country): not stated (not stated)	Group A: cefoxitin 2 g every 6 h for minimum of 4 days and continued until the woman was afebrile with improvement of symptoms for at least 48 h Group B: cefotaxime 2 g every 8 h for minimum of 4 days and continued until the woman was afebrile with improvement of symptoms for at least 48 h Group C: clindamycin 900 mg every 8 h + gentamicin at initial loading dose of 120 mg, followed by maintenance doses of 80 mg every 8 h. Subsequent maintenance doses determined by evaluating trough and peak serum aminoglycoside concentrations. Antibiotics given for minimum of 4 days, and continued until the woman was afebrile with improvement of symptoms for at least 48 h	Primary outcome: clinical failure: evaluated on a daily basis. Women who had not demonstrated signs of improvement after 48-72 h of antibiotic therapy were considered an unsuccessful result
Martens (1993) ²³	Number of women randomized: total: 295; group A: 150; group B: 145 Number of women analyzed: total: 249; group A: 128; group B: 121 Age (mean \pm SD) (years): group A: 25.9 \pm 5.8; group B: 26.0 \pm 6.8 Number of centres (country): 16 (US)	Group A: ofloxacin 400 mg PO every 12 h for 10 days Group B: cefoxitin 2 g IM + probenecid 1 g PO, followed by doxycycline 100 mg PO every 12 h for 10 days	Primary outcomes: clinical cure: complete resolution of tenderness; clinical improvement: partial resolution of tenderness without the need for additional antibiotic therapy Secondary outcomes: microbial cure of <i>Chlamydia trachomatis</i> , microbial cure of <i>Neisseria gonorrhoeae</i>
Okada (1988) ³²	Number of women randomized: total: 253; group A: 124; group B: 129 Number of women analyzed: total: 209; group A: 104; group B: 105 Age (years): only presented as frequencies within age groups Number of centres (country): 55 (Japan)	Group A: ciprofloxacin 200 mg PO + dummy cefroxadine placebo, 3 times daily for 7 consecutive days Group B: cefroxadine 250 mg PO + dummy ciprofloxacin placebo, 3 times daily for 7 consecutive days	Primary outcomes: clinical cure at 7th day of treatment: excellent: clinical marked improvement and clearance of bacteria; good: clear clinical improvement; poor: no clear clinical improvement; adverse events Secondary outcome: local tenderness around uterus

Ross (2006) ²⁶	<p>Number of women randomized: 749 Number of women analyzed: group A: 384; group B: 365 Age (mean \pm SD) (years): group A: 30.1 \pm 8.4; group B: 30.5 \pm 8.5 Number of centres (country): 13 (Denmark; Finland; France; Germany; Greece; Hungary; Italy; Lithuania; Poland; Russia; South Africa; Sweden; UK)</p>	<p>Group A: moxifloxacin 400 mg PO once daily for 14 days. Group B: ofloxacin 400 mg PO twice daily + metronidazole 500 mg PO twice daily for 14 days</p>	<p>Primary outcome: clinical cure (5-24 days post-therapy): reduction of the pelvic pain score by > 70% (McCormack score) + apyrexia (rectal/tympanic/oral temperature < 38.0 °C or axillary/cutaneous temperature < 37.5 °C) + WBC count < 10,500/mm³. Secondary outcomes: microbial cure of <i>Chlamydia trachomatis</i>; microbial cure of <i>Neisseria gonorrhoeae</i></p>
Roy (1985) ⁴⁶	<p>Number of women randomized: 46 (36 with acute PID) Number of women analyzed: group A: 19; group B: 9; group C: 9 Age (years): not stated Number of centres (country): 1 (US)</p>	<p>Group A: cefotaxime 2 g IV or IM every 8 h Group B: clindamycin 600 mg IV every 6 h + gentamicin 1.5 mg/kg lean bodyweight IV or IM every 8 h Group C: clindamycin 600 mg IV every 6 h + gentamicin 1.5 mg/kg lean bodyweight IV or IM every 8 h + penicillin G 5 million units IM every 4 h</p>	<p>Primary outcomes: clinical cure: antibiotic change was made for treatment failure from an assigned regimen based upon persistence or worsening of signs and symptoms after 48 h</p>
Roy (1990) ⁴⁷	<p>Number of women randomized: 67 Number of women analyzed: total: 67; group A: 13; group B: 14; group C: 19; group D: 21 Age (mean \pm SEM) (years): group A: 26.6 \pm 1.9; group B: 28.9 \pm 1.7; group C: 27.1 \pm 1.3; group D: 28.3 \pm 1.0 Number of centres (country): 1 (US)</p>	<p>Group A: ceftizoxime 2 g IV every 12 h + doxycycline 100 mg IV twice daily Group B: ceftizoxime 2 g IV every 6 h + doxycycline 100 mg IV twice daily Group C: ceftizoxime 2 g IV every 8 h + doxycycline 100 mg IV twice daily Group D: clindamycin 900mg IV every 8 h + gentamicin 2mg/kg loading dose followed by 1.5 mg/kg IV every 8 h with adjustments if necessary</p>	<p>Primary outcomes: clinical cure: adequate response to therapy: clinically improved and afebrile for 48 h at the time of discharge; 8-24 before discharge; no pelvic tenderness; adverse events leading to discontinuation of therapy Secondary outcomes: microbial cure of <i>Chlamydia trachomatis</i>, microbial cure of <i>Neisseria gonorrhoeae</i>, and length of hospital stay</p>
Savaris (2007) ²¹	<p>Number of women randomized: 133 Number of women analyzed: group A: 66; group B: 67 Age (mean \pm SD) (years): group A: 28.3 \pm 0.8; group B: 29.27 \pm 1.1 Number of centres (country): 1 (Brazil)</p>	<p>Group A: ceftriaxone IM 250 mg + azithromycin 1 g PO single dose and repeated after 7 days Group B: ceftriaxone IM 250 mg + doxycycline 200 mg PO for 14 days</p>	<p>Primary outcome: clinical cure defined as \geq70% reduction in the total tenderness score at day 14 compared with baseline, for both visual analogue scale and McCormack pain scale Secondary outcome: microbial cure of <i>Chlamydia trachomatis</i></p>

Sirayapiwat (2002) ⁴³	<p>Number of women randomized: 44; 22 in each group Number of women analyzed: 44; 22 in each group Age (mean \pm SD) (years): group A: 31.2 \pm 9.1; group B: 24.9 \pm 7.6 Number of centres (country): 1 (Thailand)</p>	<p>Group A: (triple therapies) received intravenous therapy of 1 gm of ampicillin every 6 hours plus 5 mg/kg. (not exceed 240 mg) of gentamicin once daily and 500 mg of Metronidazole every 8 hours</p> <p>Group B: clindamycin 600 mg IV every 8 h + gentamicin 5 mg/kg not exceeding a maximum dose of 240 mg once daily</p> <p>In both groups, parenteral therapies continued until women were afebrile for minimum of 48 h then all women received a regimen of doxycycline 100 mg PO every 12 h to complete a 14-day course</p>	<p>Primary outcomes: clinical cure; adverse events leading to discontinuation of therapy Secondary outcome: length of hospital stay. Visual analogue scale on day 3 of treatment</p>
Soper (1988) ⁴⁸	<p>Number of women randomized: 62; 31 in each group Number of women analyzed: 62; 31 in each group Age (mean \pm SD) (years): group A: 23.4 \pm 5.8; group B: 21.9 \pm 3.7 Number of centres (country): 1 (US)</p>	<p>Group A: cefoxitin 2 g IV every 6 h + doxycycline 100 mg IV every 12 h. Women discharged with doxycycline 100 mg PO twice daily to complete a 10-day course</p> <p>Group B: clindamycin 600 mg IV every 6 h + amikacin 7.5 mg/kg IV every 12 h. Women discharged with clindamycin 300 mg PO 4 times daily to complete a 10-day course</p>	<p>Primary outcome: clinical failure: persistence of fever ($>$ 38 $^{\circ}$C), elevated WBC ($>$ 11,000/mm³), moderate-severe pelvic organ tenderness despite 96 h of antibiotic therapy, need of laparotomy Secondary outcome: length of hospital stay</p>
Sweet (1985) ²⁹	<p>Number of women randomized: 60; 30 in each group Number of women analyzed: 60; 30 in each group Age (mean \pm SD) (years): group A: 24.2 \pm 5.1; group B: 24.7 \pm 7.2 Number of centres (country): 1 (US)</p>	<p>Group A: moxalactam 2 g IV every 8 h</p> <p>Group B: clindamycin 600 mg IV every 6 h + tobramycin 1.5 mg/kg every 8 h</p>	<p>Primary outcome: microbiological cure Secondary outcome: length of hospital stay</p>
Thadepalli (1991) ⁴⁵	<p>Number of women randomized: 71; group A: 35; group B: 36 Number of women analyzed: 61; group A: 31; group B: 30 Age (mean (range)) (years): group A: 27.5 (15-37); group B: 26.5 (19-36) Number of centres (country): not stated (not stated)</p>	<p>Group A: ciprofloxacin 300 mg IV twice daily for \geq 3 days followed by ciprofloxacin 500 mg PO twice daily for about 1 week</p> <p>Group B: clindamycin 600 mg IV every 6 h + gentamicin 80 mg IV every 8 h, administered separately. Clindamycin by IV route for 3 days, followed by oral administration for about 1 week. Gentamicin dose adjusted based on serum creatinine and gentamicin levels</p>	<p>Primary outcomes: clinical cure: when there was resolution or clearing (or both) of signs of infection as evidenced by defervescence, reversal of leukocytosis, and abatement of abdominal pain and cervical motion tenderness; adverse events leading to discontinuation of therapy Secondary outcome: length of hospital stay</p>
Tison (1988) ²⁷	<p>Number of women randomized: 40; 20 in each group Number of women analyzed: 40; 20 in each group</p>	<p>Group A: amoxicillin-clavulanate 1 g slow IV perfusion every 8 h. PO once temperature normal for 48 h, 4 tablets of 500 mg, 2 at a time as long as hospitalized. After discharge, treatment continued with 3 tablets daily of amoxicillin-clavulanate for 3 weeks</p>	<p>Effectiveness: cure: absence of pelvic pain on examination and referred by women, normal body temperature at discharge and 3 weeks later</p>

	Age (mean (range)) (years): group A: 27.5 (17-39); group B: 22.5 (17-34) Number of centres (country): 1 (France)	Group B: penicillinG10million units/24 h IV, followed by penicillinVPO+ gentamicin 160 mg/24 h IM for 10 days + metronidazole 500 mg IV 3 times daily followed by metronidazole 500 mg PO 3 times daily Duration of treatment unclear.	Adverse events: any antibiotic-related adverse event leading to discontinuation of therapy during hospital stay and after 30 days
Walters (1990) ³²	Number of women randomized: 147; unclear how many in each group Number of women analyzed: group A: 63; group B: 67. PP analysis only Age (mean \pm SD) (years): total age not stated; group A: 25.4 \pm 7.7; group B: 24.5 \pm 7.8 Number of centres (country): 1 (US)	Group A: gentamicin 2.0 mg/kg IV as loading dose, then 1.5 mg/kg IV every 8 h + clindamycin 900 mg IV every 8 h for minimum 4 days. After discharge, clindamycin 450 mg PO every 6 h for total 14 days Group B: cefoxitin 2 g IV every 6 h + doxycycline 100 mg IV every 12 h for minimum 4 days. After discharge, doxycycline 100 mg PO every 12 h for total 14 days	Primary outcomes: clinical cure defined as oral temperature < 38 °C for 48 h, resolution of pain and tenderness, and no increase in the size of any pelvic mass after 21 days of initiation of treatment Secondary outcomes: microbiological clearance of <i>Chlamydia trachomatis</i> ; microbiological clearance of <i>Neisseria gonorrhoeae</i> after 21 days of initiation of treatment
Wendel (1991) ²²	Number of women randomized: 96; unclear how many to each group Number of women analyzed: group A: 35; group B: 37. PP analysis only Age (mean) (years): total age not stated; group A: 24.7; group B: 23.3 Number of centres (country): 1 (US)	Group A: cefoxitin 2 g IM + probenecid 1 g PO followed by doxycycline 100 mg PO every 12 h for 10 days Group B: ofloxacin 400 mg PO every 12 h for 10 days	Primary outcomes: clinical cure defined as complete resolution of tenderness (> 65% decrease in clinical score); adverse events leading to discontinuation of therapy Secondary outcomes: microbiological clearance of <i>Chlamydia trachomatis</i> ; microbiological clearance of <i>Neisseria gonorrhoeae</i>

ESR: erythrocyte sedimentation rate; h: hour; IM: intramuscular; ITT: intention to treat; IUD: intrauterine device; IV: intravenous; n: number of women; PID: pelvic inflammatory disease; PO: per os; PP: per protocol; SD: standard deviation; WBC: white blood cell