

# Efficacy and safety of Kanglaite injection combined with first-line platinum-based chemotherapy in patients with advanced NSCLC: a systematic review and meta-analysis of 32 RCTs

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**Background:** As a Chinese medicine injections, Kanglaite injection (KLT) is a complementary or alternative therapy for first-line platinum-based chemotherapy. However, the effect that certain factors, including the dose of KLT, chemotherapy cycles, evaluation criteria, or supportive treatment, have on the efficacy of the objective response rate (ORR), median survival time (MST), and adverse reactions is still unknown.

**Methods:** Eight databases were systematically searched from the inception dates to December 1, 2019, using the keywords Kanglaite, chemotherapy, and non small cell lung carcinoma to identify randomized clinical trials (RCTs). Analyses were performed using Review Manager 5.3 and Stata 15.1.

**Results:** There were 32 randomized controlled trials, involving 2,577 participants, that fulfilled the inclusion criteria. Compared with first-line platinum-based chemotherapy alone, KLT combined with chemotherapy could increase the ORR [risk ratio (RR), 1.41 (95% CI: 1.28 to 1.56); absolute risk difference (ARD), 0.13 (95% CI: 0.1 to 0.17)], decrease the risk ratio of adverse reactions [nausea and vomiting: RR, 0.58 (95% CI: 0.42 to 0.81); ARD, -0.17 (95% CI: -0.26 to -0.08); leukopenia: RR, 0.61 (95% CI: 0.44 to 0.86); ARD, -0.16 (95% CI: -0.24 to -0.08)], prolong MST, and increase disease control rate and Karnofsky performance status. According to the subgroup analyses, KLT combined with cisplatin or paraplatin plus paclitaxel (TP) failed to demonstrate a significant association with the ORR. And when lacking the use of supportive treatment, this combination would not decrease the RR of both adverse reactions compared with chemotherapy alone.

**Conclusions:** KLT plus first-line platinum-based chemotherapy, except when chemotherapy regimens were TP, increased efficacy and quality of life in patients with advanced NSCLC. We are unsure whether this combination offers a low risk of adverse reactions. Additional high-quality RCTs are warranted to assess the effects of the combined therapy further.

**Keywords:** Kanglaite injection; first-line chemotherapy; non-small cell lung carcinoma (NSCLC); efficacy; metaanalysis

Submitted Mar 11, 2020. Accepted for publication Jun 05, 2020. doi: 10.21037/apm-20-616 View this article at: http://dx.doi.org/10.21037/apm-20-616

#### Introduction

Lung cancer is the most commonly diagnosed cancer. According to global cancer statistics, lung cancer was the leading cancer-related death in 2018 (1), with 80% of these cases being non-small cell lung carcinomas (NSCLC). Even with this well-known mortality, over 50% of NSCLC present with advanced local invasion and metastasis during hospital admission diagnosis, meaning they have missed the opportunity for surgical intervention. Despite the promising emergence of molecular targeted therapy and immunotherapy, platinum-based chemotherapy is still the cornerstone of NSCLC treatment, especially for advanced stages III and IV of the disease (2). Firstline platinum-based chemotherapy, including cisplatin or paraplatin plus vinorelbine, paclitaxel, gemcitabine, docetaxel, or pemetrexed (3), is widely used in advanced NSCLC. However, adverse reactions, including nausea and leukopenia, are frequently reported (4,5).

In traditional Chinese medicine (TCM) basic theory, lung cancer is nearly equivalent to the domains of "mass" or "phlegm and dampness", which is acknowledged to be one of the basic pathogeneses. Kanglaite injection (KLT) (Z10970091, China Food and Drug Administration) is extracted from the seeds of the Chinese medicinal herb (CMH) Coix lacryma-jobi, whose anticancer effect thought treating the dampness with bland and treat "mass" in traditional Chinese medicine theory. Past studies have suggested that KLT is a micro-emulsion for intravenous use that demonstrates antitumor efficacy, improves the quality of life (QOL), and reduces toxicity (6,7). However, the outcome of its combination with different chemotherapy regimens and the long-term synergistic efficacy is still unclear. Moreover, CMH is often considered to have serious adverse reactions (8), and the Chinese government has announced a post-marketing review of TCM injections in the following 5 to 10 years (9). Finally, the exact effect and survival rate after the application of KLT are also a concern.

This systematic review and meta-analysis were performed to compare KLT plus first-line platinum-based chemotherapy with first-line platinum-based chemotherapy alone in patients with advanced NSCLC by the using tumor response and adverse reactions as outcome measures. We present the following article following the PRISMA 2009 reporting checklist (available at http://dx.doi.org/10.21037/ apm-20-616).

#### Methods

This article follows the Preferred Reporting Items for Systematic Reviews and Meta-Analyses guidelines (PRISMA guidelines), and the study is registered with PROSPERO (CRD 42019142414). As a systematic review and metaanalysis, Ethical approval was not required as materials of this study had been published.

#### Data sources

Two reviewers (Juan Li and Hong-Zheng Li) independently searched for and extracted information from randomized clinical trials (RCTs) related to KLT-assisted treatment of NSCLC. RCTs were searched for in Chinese and English databases, including the PubMed, Cochrane Library, EMBASE, Web of Science (ISI), Chinese National Knowledge Infrastructure (CNKI), Chinese Scientific Journals Full-Text Database (VIP), CBM, and Wanfang databases. The searches were restricted to original publications from the time of establishment to December 1, 2019. A combination of the following keywords was used: "lung cancer", "lung carcinoma", "non-small cell lung cancer", "non-small lung carcinoma", "NSCLC", "Kanglaite", "KLT", and "Coix Seed Oil". All retrievals were implemented using the Medical Subject Headings (MeSH) and free word. Finally, all related systematic reviews (SRs) and meta-analyses were evaluated, and studies meeting the inclusion criteria were selected from the references. As an example, the electronic strategy for PubMed can be seen in Figure S1.

# Search strategies and selection criteria

Trials were selected on the following inclusion criteria: (I) the trial was an RCT. (II) The patients were diagnosed with NSCLC stages III to IV, according to histopathological and cytological diagnostic criteria. (III) The experimental group had undergone KLT (Z10970091, China Food and Drug Administration) plus first-line platinum-based chemotherapy, and the control group had undergone first-line platinum-based chemotherapy refers to cisplatin or paraplatin plus vinorelbine, paclitaxel, gemcitabine, docetaxel or pemetrexed (NP, TP, GP, DP, and AP, respectively). (IV) Patients did not receive any radiotherapy, other chemotherapy, or Chinese herbs during this study. (V) The

outcome needed to include at least an objective response rate (ORR) or adverse reactions (nausea and vomiting, leukopenia). Exclusion criteria were (I) duplicates (797 studies); (II) unrelated studies including other treatments (56 studies); (III) non-RCTs including case-control studies and series case reports (23 studies); (IV) abstracts and reviews without specific data and unrelated SRs (59 studies), and (V) studies with no ORR or adverse reaction (nausea and vomiting) data (17 studies).

#### Data extraction and quality assessment

Two researchers (Juan Li and Guang-Hui Zhu) independently extracted the following information from each study: the lead author; the publication time; the demographic characteristics; the sample size; the usage of KLT and the types of first-line platinum-based chemotherapy; the evaluation criteria of clinical efficacy; and whether supportive treatment including anti-nausea drugs, granulocyte colony-stimulating factor (G-CSF) were administered. Furthermore, outcomes including the ORR, leukopenia, nausea and vomiting, median survival time (MST), disease control rate (DCR), and Karnofsky performance status (KPS) were examined. The data were obtained directly from the articles. A third reviewer resolved any disagreements (Jie Li).

The methodological quality of the included RCTs was assessed independently by two researchers (Hong-Zheng Li and Guang-Hui Zhu) on the Cochrane risk-of-bias criteria (10) using the following parameters to evaluate the bias risk: random sequence generation (selection bias), allocation concealment (selection bias), blinding of participants and personnel (performance bias), blinding of outcome assessment (detection bias), incomplete outcome data (attrition bias), selective reporting (reporting bias), and other bias (whether the baseline is comparable). Subsequently, the trials were assessed and categorized into three levels: low risk (all items were "yes"), high risk (at least one item was "no"), and unclear risk (at least one item was "unclear").

#### Main outcomes

We measured the tumor response using the ORR. According to the World Health Organization (WHO) guidelines for solid tumor responses (11) or Response Evaluation Criteria in Solid Tumors (RECIST) (12), indicators were complete response (CR), partial response (PR), stable disease (SD), progressive disease (PD), with ORR being equal to CR plus PR. Adverse reactions (adverse drug events or adverse drug reactions) were pooled, including nausea and vomiting, and leukopenia.

# Secondary outcomes

The long-term synergistic efficacy of this combination was considered MST. Also, the secondary outcomes included DCR and QOL. QOL was considered improved if the KPS score increased by 10 points or higher after treatment (13). DCR was calculated as CR plus PR and SD.

# Statistical analysis

Two reviewers performed the meta-analysis (Juan Li and Guang-Hui Zhu) using Review Manager 5.3 (The Cochrane Collaboration, Oxford, UK) and Stata 15.1. The relative risk (RR) and 95% confidence intervals (CI) were calculated. Statistical heterogeneity of the results across trials was assessed by a Chi-square based Q-statistic test, and the consistency was calculated by I<sup>2</sup>. If the homogeneity (P $\geq$ 0.1, I<sup>2</sup> $\leq$ 50%) was not rejected, the fixed-effects model (FEM) was used to calculate the summary RR and the 95% CI. Alternatively, the results were calculated by the randomeffects model (REM). We performed a subgroup analysis according to different doses of KLT, types of first-line platinum-based chemotherapy, the cycle of chemotherapy, and evaluation criteria, which revealed their influence on tumor responses.

Furthermore, we performed a subgroup analysis regarding using supportive treatment for adverse reactions. Publication bias was evaluated using funnel plots if more than 10 included studies were included. Subgroup analyses were performed following the doses of KLT, the type, and cycle of chemotherapy, supportive treatment, and evaluation criteria to reveal the clinical heterogeneity and its influence on the endpoint.

# Results

The literature search identified 188 studies and excluded 68 after careful screening of titles and abstracts. The full text of the 120 remaining studies was assessed for eligibility, and 88 were excluded because they were reviews, did not contain eligible comparators, did not report outcomes of interest, were case series, or for other reasons (*Figure 1*). Finally, 32 articles comprising 2,577 patients met the inclusion criteria



Figure 1 Articles retrieved and assessed for eligibility.

and were retrieved for quantitative synthesis, with all the records being studied in China (*Table 1*).

# Characteristics of eligible studies

The experimental group comprised 291 cases of KLT plus first-line platinum-based chemotherapy, while the control group comprised 1,286 cases of first-line platinum-based chemotherapy alone. There were 1,436, and 989 males and females included respectively, with age ranging between 32 and 80 years. The dosage of KLT was 60 to 300 mL/day, and the treatment time was 3 to 4 weeks/cycle, with 1 to 4 cycles of intravenous injection. According to the WHO guidelines for solid tumor responses (11) or RECIST (12), tumor responses were evaluated in 32 studies (14-29) involving 2,577 patients (30-45). Nausea and vomiting were evaluated in 9 studies (14,19,22,29,37,38,42,43,45) involving 692 patients, and leukopenia was evaluated in 11 studies (17,19,21,22,26,29,37-39,42,45) involving 901 patients. According to the WHO standards (11) or National Cancer Institute Common Toxicity Criteria (NCI-CTC) (46), 4 studies for nausea and 5 studies for leukopenia were included.

## Methodological quality

All studies mentioned randomization, with only 17 studies (14,16,17,20,22,23,26-28,30,33-36,39,43,45) reporting the details of the randomized methods, and none reporting the details of concealed allocations. One study (38) revealed the blinding methods but not the details regarding the blinding of patients or assessors. Additionally, one study (33) revealed that all the participants were aware of the treatment in advance, which did not affect the outcome.

Fourteen participants withdrew from three studies (27,36,41), with five presenting acute/subacute toxicity, five participants were treated with another treatment, and four participants withdrew for personal or economic reasons. Furthermore, seven studies lacked outcome data (19,20,24,34,35,40,44). Three studies selectively reported acute/subacute toxicity (34,40,44), with one reporting KPS (33). The methodological bias risk of all included studies is presented in *Table 2* and *Figure 2*.

# ORR

The reporting of ORR occurred in 32 studies (14-29)

Table 1 Characteristics of included studies

|                    | N     | ISCLC (III- | -IV)           | Inventions |           |           |           |           |                         | Ou      | tcome     |
|--------------------|-------|-------------|----------------|------------|-----------|-----------|-----------|-----------|-------------------------|---------|-----------|
| First author, year | F/0   |             | <b>A</b> = = = | Treat      | ment      | Osistinal | Scale (A) | Scale (B) | Supportive<br>treatment | Main    | Secondary |
|                    | E/C   | IVI/F       | Age            | E          | KLT (D/C) | Control   |           |           | troutmont               | outcome | outcomes  |
| Bao 2019, (14)     | 31/31 | 38/24       | 39–72          | GP + KLT   | 200 mL/2  | GP        | WHO       | Unclear   | No                      | 53      | 5         |
| Chen 2018, (15)    | 30/30 | Unclear     | 35–65          | GP + KLT   | 200 mL/1  | GP        | RECIST    | Unclear   | Yes                     |         | 5         |
| Chen 2016, (16)    | 44/44 | 47/41       | 55–78          | GP + KLT   | ?/4       | GP        | RECIST    | Unclear   | No                      |         | 5         |
| Chen 2018, (17)    | 51/51 | 59/43       | 57–79          | GP + KLT   | 200 mL/4  | GP        | WHO       | Unclear   | No                      | 52      | 5         |
| Chen 2003, (18)    | 28/27 | 33/22       | 33–80          | NP + KLT   | 200 mL/2  | NP        | WHO       | Unclear   | No                      |         | 56        |
| Guan 2009, (19)    | 12/12 | 11/12       | 36–72          | GP + KLT   | 300 mL/2  | GP        | WHO       | NCI-CTC   | Yes                     | 423     | 45        |
| He 2017, (20)      | 54/54 | 81/27       | 38–80          | DP + KLT   | 100 mL/3  | DP        | WHO       | Unclear   | Yes                     |         |           |
| Huang 2010, (21)   | 35/35 | 44/26       | 59–78          | GP + KLT   | 200 mL/2  | GP        | WHO       | WHO       | No                      | 52      | 56        |
| Jia 2018, (22)     | 31/31 | 34/28       | 43–74          | DP + KLT   | 200 mL/2  | DP        | WHO       | Unclear   | Yes                     | 523     | 5         |
| Li 2017, (23)      | 41/41 | 43/39       | 55–75          | GP + KLT   | ?/4       | GP        | RECIST    | Unclear   | No                      |         | 5         |
| Li 2012, (24)      | 38/40 | 49/29       | 70–77          | TP + KLT   | 100 mL/3  | TP        | RECIST    | Unclear   | No                      |         | 56        |
| Li 2016, (25)      | 39/39 | 43/35       | 35–72          | NP + KLT   | 200 mL/2  | NP        | WHO       | WHO       | Yes                     |         | 5         |
| Liang 2018, (26)   | 40/40 | 44/36       | Unclear        | NP + KLT   | 100 mL/2  | NP        | WHO       | Unclear   | Yes                     | 52      | 5         |
| Liang 2014, (27)   | 23/20 | Unclear     | 60–75          | GP + KLT   | 100 mL/2  | GP        | WHO       | Unclear   | No                      |         | 56        |
| Liu 2019, (28)     | 63/63 | 79/47       | 50–77          | GP + KLT   | 200 mL/2  | GP        | RECIST    | Unclear   | Yes                     | 1       | 456       |
| Liu 2011, (29)     | 35/35 | 44/26       | 59–74          | GP + KLT   | 200 mL/2  | GP        | RECIST    | WHO       | Yes                     | 523     | 56        |
| Liu 2016, (30)     | 55/55 | 62/48       | 45–79          | NP + KLT   | 200 mL/4  | NP        | RECIST    | Unclear   | No                      |         | 5         |
| Liu 2015, (31)     | 43/43 | 55/31       | 42–74          | GP + KLT   | 200 mL/4  | GP        | RECIST    | SFDA      | Yes                     |         | 56        |
| Long 2017, (32)    | 42/40 | 52/30       | 47–70          | GP + KLT   | 200 mL/3  | GP        | RECIST    | WHO       | No                      |         | 5         |
| Mu 2018, (33)      | 47/47 | 56/38       | 45–76          | DC + KLT   | 200 mL/2  | DP        | RECIST    | Unclear   | No                      |         | 5         |
| Sun 2012, (34)     | 35/35 | 41/29       | 37–75          | GP + KLT   | 200 mL/4  | GP        | WHO       | WHO       | No                      |         | 5         |
| Wang 2018, (35)    | 42/42 | 45/39       | 55–78          | DP + KLT   | 100 mL/4  | DP        | RECIST    | Unclear   | Yes                     |         |           |
| Wang 2016, (36)    | 24/25 | 39/13       | 55–70          | AP + KLT   | 200 mL/2  | AP        | RECIST    | WHO       | Yes                     |         | 5         |
| Wang 2014, (37)    | 43/43 | 58/28       | 43–79          | GP + KLT   | 200 mL/1  | GP        | RECIST    | WHO       | Yes                     | 523     | 56        |
| Wang 2017, (38)    | 36/36 | 32/40       | Unclear        | GP + KLT   | 60 mL/4   | GP        | WHO       | Unclear   | No                      | 523     | 56        |
| Yan 2018, (39)     | 49/49 | 63/35       | 38–76          | GP + KLT   | 200 mL/4  | GP        | RECIST    | Unclear   | Yes                     | 52      | 5         |
| Yang 2016, (40)    | 35/35 | 33/37       | Unclear        | TP + KLT   | 100 mL/3  | TP        | RECIST    | Unclear   | No                      |         | 5         |
| Yang 2003, (41)    | 25/26 | Unclear     | 36–68          | NP + KLT   | 200 mL/3  | NP        | WHO       | Unclear   | No                      |         | 56        |
| Yao 2017, (42)     | 70/67 | 78/59       | Unclear        | GP + KLT   | 200 mL/2  | GP        | RECIST    | WHO       | No                      | 523     | 5         |
| Ye 2017, (43)      | 40/40 | 54/26       | 55–74          | GP + KLT   | 200 mL/2  | GP        | RECIST    | Unclear   | No                      | 53      | 5         |
| Yu 2015, (44)      | 60/60 | 67/53       | Unclear        | TP + KLT   | 100 mL/3  | TP        | WHO       | Unclear   | Yes                     |         | 5         |
| Zhang 2019, (45)   | 50/50 | 52/48       | Unclear        | GP + KLT   | ?/3       | GP        | RECIST    | Unclear   | No                      | 423     | 45        |

NSCLC, non-small cell lung cancer; E/C, experimental group (Kanglaite injection plus first-line platinum-based chemotherapy)/control group (first-line platinum-based chemotherapy); M/F, male/female; KLT (D/C), dose/cycles; GP, cisplatin or paraplatin and gemcitabine; NP, cisplatin or paraplatin and vinorelbine; TP, cisplatin or paraplatin and paclitaxel; DP, cisplatin or paraplatin and docetaxel; AP, cisplatin or paraplatin and pemetrexed; scale. A, evaluation criteria of tumor response; scale. B, evaluation criteria of adverse reactions; RECIST, response evaluation criteria in solid tumors; NCI-CTC, National Cancer Institute Common Toxicity Criteria. ①, ORR = CR + PR; ②, leukopenia; ③, nausea and vomiting; ④, median survival time; ⑤, DCR = CR + PR + SD; ⑥, Karnofsky performance status (KPS).

Table 2 Risk of bias summary: the review authors' judgments about each risk-of-bias item for each included randomized control trial

|                    | Dandam agguanag      | Allegation  | Blindi                     | ng                    | Incomplete   | Coloctivo |            |  |
|--------------------|----------------------|-------------|----------------------------|-----------------------|--------------|-----------|------------|--|
| First author, year | generation           | concealment | Participants and personnel | Outcome<br>assessment | outcome data | reporting | Other bias |  |
| Bao 2019           | Simple randomization | Unclear     | Unclear                    | Unclear               | Complete     | Low risk  | High risk  |  |
| Chen 2018          | Random draw          | Unclear     | Unclear                    | Unclear               | Complete     | Low risk  | High risk  |  |
| Chen 2016          | Simple randomization | Unclear     | Unclear                    | Unclear               | Complete     | Low risk  | Low risk   |  |
| Chen 2018          | Simple randomization | Unclear     | Unclear                    | Unclear               | Complete     | Low risk  | High risk  |  |
| Chen 2003          | Unclear              | Unclear     | Unclear                    | Unclear               | Complete     | Low risk  | Unclear    |  |
| Guan 2009          | Unclear              | Unclear     | Unclear                    | Unclear               | Incomplete   | Low risk  | High risk  |  |
| He 2017            | Simple randomization | Unclear     | Unclear                    | Unclear               | Incomplete   | Low risk  | Low risk   |  |
| Huang 2010         | Unclear              | Unclear     | Unclear                    | Unclear               | Complete     | Low risk  | High risk  |  |
| Jia 2018           | Simple randomization | Unclear     | Unclear                    | Unclear               | Complete     | Low risk  | Low risk   |  |
| Li 2017            | Simple randomization | Unclear     | Unclear                    | Unclear               | Complete     | Low risk  | High risk  |  |
| Li 2012            | Unclear              | Unclear     | Unclear                    | Unclear               | Incomplete   | High risk | High risk  |  |
| Li 2016            | Unclear              | Unclear     | Unclear                    | Unclear               | Complete     | Low risk  | Unclear    |  |
| Liang 2018         | Simple randomization | Unclear     | Unclear                    | Unclear               | Complete     | Low risk  | High risk  |  |
| Liang 2014         | Simple randomization | Unclear     | Unclear                    | Unclear               | Incomplete   | Low risk  | High risk  |  |
| Liu 2019           | Simple randomization | Unclear     | Unclear                    | Unclear               | Complete     | Low risk  | High risk  |  |
| Liu 2011           | Unclear              | Unclear     | Unclear                    | Unclear               | Complete     | Low risk  | High risk  |  |
| Liu 2016           | Simple randomization | Unclear     | Unclear                    | Unclear               | Complete     | Low risk  | Low risk   |  |
| Liu 2015           | Unclear              | Unclear     | Unclear                    | Unclear               | Complete     | Low risk  | High risk  |  |
| Long 2017          | Unclear              | Unclear     | Unclear                    | Unclear               | Complete     | Low risk  | High risk  |  |
| Mu 2018            | Simple randomization | Unclear     | Unclear                    | Unclear               | Complete     | Low risk  | High risk  |  |
| Sun 2012           | Simple randomization | Unclear     | Unclear                    | Unclear               | Incomplete   | High risk | Unclear    |  |
| Wang 2018          | Simple randomization | Unclear     | Unclear                    | Unclear               | Incomplete   | Low risk  | Unclear    |  |
| Wang 2016          | Simple randomization | Unclear     | Unclear                    | Unclear               | Incomplete   | Low risk  | Unclear    |  |
| Wang 2014          | Unclear              | Unclear     | Unclear                    | Unclear               | Complete     | Low risk  | Low risk   |  |
| Wang 2017          | Unclear              | Unclear     | Yes                        | Unclear               | Complete     | Low risk  | High risk  |  |
| Yan 2018           | Simple randomization | Unclear     | Unclear                    | Unclear               | Complete     | Low risk  | High risk  |  |
| Yang 2016          | Unclear              | Unclear     | Unclear                    | Unclear               | Incomplete   | High risk | High risk  |  |
| Yang 2003          | Unclear              | Unclear     | Unclear                    | Unclear               | Incomplete   | Low risk  | High risk  |  |
| Yao 2017           | Unclear              | Unclear     | Unclear                    | Unclear               | Complete     | Low risk  | Low risk   |  |
| Ye 2017            | Simple randomization | Unclear     | Unclear                    | Unclear               | Complete     | Low risk  | High risk  |  |
| Yu 2015            | Unclear              | Unclear     | Unclear                    | Unclear               | Incomplete   | High risk | High risk  |  |
| Zhang 2019         | Simple randomization | Unclear     | Unclear                    | Unclear               | Complete     | Low risk  | High risk  |  |



**Figure 2** Risk of methodological bias. (A) Risk of bias summary: review authors' judgments about each risk of bias item for each included study. (B) Risk of bias graph: review authors' judgments about each risk of bias item presented as percentages across all included studies.

comprising 2,577 cases (30-45) (*Figure 3*). Pearson's Chisquare test and I<sup>2</sup> test indicated the absence of statistical heterogeneity among studies (I<sup>2</sup>=0%). The ORR in the meta-analysis demonstrated statistical differences between KLT plus first-line platinum-based chemotherapy and firstline platinum-based chemotherapy alone (RR =1.41, 95% CI: 1.28 to 1.56, ARD =0.13, 95% CI: 0.1. to 0.17, and P<0.00001) by FEM.

# Adverse reactions

A high heterogeneity was observed among studies regarding nausea and vomiting ( $I^2=64\%$ ) (*Figure 4*) and leukopenia ( $I^2=77\%$ ) (*Figure 5*). The meta-analysis revealed that KLT plus first-line platinum-based chemotherapy involved a lower risk of nausea and vomiting (RR =0.58, 95% CI: 0.42 to 0.81, ARD =-0.17, 95% CI: -0.26 to -0.08, and P=0.001) and leukopenia (RR =0.61, 95% CI: 0.44 to 0.86, ARD =-0.16, 95% CI: -0.24 to -0.08, and P=0.004) than the first-line platinum-based chemotherapy alone using REM. Furthermore, all differences were statistically significant.

#### Effects on secondary outcomes

For secondary outcomes, 3 studies with 243 cases (19,28,45) mentioned the average value of MST (*Figure 6, Table 3*), while 10 studies with 899 cases (18,21,24,27,29-31,37,38,41) reported the KPS scale evaluated by QOL (*Figure 7*). DCR was reported in 31 studies with 2,493 cases (*Figure 8*). Minimal heterogeneity was observed among studies in MST ( $I^2$ =0%), KPS ( $I^2$ =0%), and DCR ( $I^2$ =2%). Meta-analysis demonstrated that MST (HR =0.37, 95% CI: 0.12 to 0.62), QOL (RR =1.82, 95% CI: 1.51 to 2.19, ARD =0.23, 95% CI: 0.16 to 0.29), and DCR (RR =1.16, 95% CI: 1.11 to 1.22, ARD =0.11, 95% CI: 0.08 to 0.15) indicated statistical differences between the two groups by FEM.

# Subgroup analysis

#### Subgroup analysis of ORR

Subgroup analysis was performed to reveal the influence of different doses, types of first-line platinum-based chemotherapy, cycles of chemotherapy, and evaluation

| Study or Subgroup   | Events  | Total   | Events   | Total   | Weight   | M-H, Fixed, 95% Cl   | M-H, Fixed, 95% Cl  |
|---|---|---|--|---|--|--|---|
| Bao H 2019  | 25  | 31  | 17   | 31  | 4,1%   | 1,47 [1,02, 2.11]  |   |
| Chen C-2018   | 16  | 30  | 11   | 30  | 2.6%   | 1.45 (0.82, 2.59)  |   |
| Chen W 2016   | 17  | 44  | 15   | 44  | 3.6%   | 1.13 [0.65, 1.97]  |   |
| Chen Y 2018   | 22  | 51  | 18   | 51  | 4.3%   | 1.22 [0.75, 1.99]  |   |
| Chen YZ 2003  | 13  | 28  | 9  | 27  | 2,2%   | 1.39 [0.72, 2.71]  |   |
| Guan XO 2009  | 5   | 12  | 4  | 12  | 1.0%   | 1.25 (0.44, 3.55)  |   |
| He LT 2017  | 22  | 54  | 11   | 54  | 2,6%   | 2.00 [1.08, 3.71]  |   |
| Huang ZB 2010   | 19  | 35  | 13   | 35  | 3.1%   | 1.46 [0.86, 2.48]  |   |
| Jia JN 2018   | 15  | 31  | 10   | 31  | 2.4%   | 1.50 [0.80, 2.81]  |   |
| LI HY 2017  | 16  | 41  | 14   | 41  | 3.4%   | 1.14 [0.65, 2.02]  |   |
| LIL 2012  | 8   | 38  | 9  | 40  | 2.1%   | 0.94 (0.40, 2,17)  |   |
| LI XD 2016  | 24  | 39  | 14   | 39  | 3.4%   | 1.71 [1.05, 2.79]  |   |
| Liang J 2018  | 23  | 40  | 10   | 40  | 2.4%   | 2.30 [1.26, 4.19]  |   |
| Liang SG 2014   | 10  | 23  | 8  | 20  | 2.1%   | 1.09 [0.53, 2.21]  |   |
| Liu F 2019  | 40  | 63  | 26   | 63  | 6.3%   | 1.54 [1.09, 2.18]  |   |
| Liu JQ 2011   | 15  | 35  | 12   | 35  | 2,9%   | 1.25 (0.69, 2.27)  |   |
| Liu XH 2016   | 27  | 55  | 17   | 55  | 4.1%   | 1.59 [0.98, 2.56]  |   |
| Liu Y 2015  | 18  | 43  | 14   | 43  | 3.4%   | 1.29 [0.74, 2.24]  |   |
| Long SG 2017  | 17  | 42  | 15   | 40  | 3.7%   | 1.08 [0.63, 1.86]  |   |
| Mu Q 2018   | 24  | 47  | 14   | 47  | 3.4%   | 1.71 [1.02, 2.89]  |   |
| Sun SQ 2012   | 15  | 35  | 11   | 35  | 2.6%   | 1.36 [0.73, 2.54]  |   |
| Wang CY 2018  | 18  | 42  | 17   | 42  | 4,1%   | 1.06 [0.64, 1.76]  |   |
| vvang HY 2016   | 10  | 24  | 8  | 25  | 1.9%   | 1.30 (0.62, 2.73)  |   |
| Wang L 2014   | 19  | 43  | 12   | 43  | 2,9%   | 1.58 [0.88, 2.85]  |   |
| wang Y 2017   | 26  | 36  | 12   | 36  | 2.9%   | 2.17 [1.31, 3.59]  |   |
| Yan QH 2018   | 23  | 49  | 21   | 49  | 5.0%   | 1.10 [0.71, 1.70]  |   |
| Yang L 2016   | 13  | 35  | 10   | 35  | Z.4%   | 1.30 [0.66, 2.56]  |   |
| Yang SJ 2003  | 9   | 25  | 1  | 26  | 1.7%   | 1.34 [0.59, 3.04]  |   |
| Ya0 J 2017  | 19  | 70  | 15   | 67  | 3.7%   | 1.21 [0.67, 2.18]  |   |
| 18 CY 2019  | 22  | 40  | 41   | 40  | 5,0%   | 1.05 (0.70, 1.57)  |   |
| 70 1 2015<br>7bobb MM 2010  | 15  | 50  | 14   | 60  | 3,4 %  | 2.50 (1.06, 5.02)  |   |
| znang mm 2015   | -14   | 50  | 0  | 50  | 1.4 %  | 2.50 (1.00, 5.52)  |   |
| Total (95% CI)  |   | 1291  |  | 1286  | 100.0%   | 1.41 [1.28, 1.56]  | •   |
|   |   |   | 11.0   |   |  |  |   |
| Total events<br>Heterogeneity: Chi <sup>2</sup> =<br>Test for overall effect  | 588<br>19.51, df  | = 31 (P   | 415<br>= 0.95); I <sup>e</sup>   | = 0%  |  |  | 0.01 0.1 1 10 100   |
| Total events<br>Heterogeneity: Chi <sup>a</sup> =<br>Test for overall effect:   | 588<br>19.51, df<br>Z = 6.91 (<br>Kangla  | = 31 (P<br>P < 0.0<br>lite  | 415<br>= 0.95); I <sup>2</sup><br>0001)<br>Chemoth   | erapy   | Weight   | Risk Difference  | 0.01 0.1 10 100<br>Favours (Kanglaite) Favours (Chemotherapy)<br>Risk Difference<br>M B Elverd 95% CL   |
| Total events<br>Heterogeneity: Chi <sup>2</sup> =<br>Test for overall effect:<br>Study or Subgroup  | 588<br>19.51, df<br>Z = 6.91 (<br>Kangla<br>Events<br>25  | = 31 (P<br>(P < 0.0<br>nite<br>Total  | 415<br>= 0.95); )*<br>00001)<br>Chemoth<br>Events  | erapy<br>Total  | Weight   | Risk Difference<br>M-H, Fixed, 95% CI  | 0.01 0.1 10 100<br>Favours (Kanglaite) Favours (Chernotherapy)<br>Risk Difference<br>M-H, Fixed, 95% Cl |
| Total events<br>Heterogeneity: Chi <sup>2</sup> =<br>Test for overall effect:<br>Study or Subgroup<br>Bao H 2019<br>Chen C 2018   | 588<br>19.51, df<br>Z = 6.91<br>Kangla<br>Events<br>25<br>16  | = 31 (P<br>P < 0.0<br>nite<br><u>Total</u><br>31<br>30  | 415<br>= 0.95);   <sup>2</sup><br>0001)<br>Chemoth<br>Events<br>17<br>11   | erapy<br>Total<br>31  | Weight<br>2.4%<br>2.3%   | Risk Difference<br>M-H, Fixed, 95% CI<br>0.26 (0.03, 0.48)<br>0.17 Job 08: 0.411   | 0.01 0.1 10 100<br>Favours (Kanglaite) Favours (Chemotherapy)<br>Risk Difference<br>M-H, Fixed, 95% Cl  |
| Total events<br>Heterogeneity: Chi <sup>2</sup> =<br>Test for overall effect:<br>Study or Subgroup<br>Bao H 2019<br>Chen C 2018<br>Chen W 2018  | 588<br>19.51, df<br>Z = 6.91 (<br>Kangla<br>Events<br>25<br>16<br>17  | = 31 (P<br>P < 0.0<br>nite<br><u>Total</u><br>31<br>30<br>44  | 415<br>= 0.95);   <sup>2</sup><br>00001)<br>Chemothe<br>Events<br>17<br>11<br>15   | = 0%<br>erapy<br><u>Total</u><br>31<br>30<br>44   | Weight<br>2.4%<br>2.3%<br>3.4%   | Risk Difference<br><u>M-H, Fixed, 95% Cl</u><br>0.26 (0.03, 0.48)<br>0.17 (-0.08, 0.41)<br>0.05 I-0.16 0.25)   | 0.01 0.1 10 100<br>Favours (Kanglaite) Favours (Chemotherapy)<br>Risk Difference<br>M-H, Fixed, 95% Cl  |
| Total events<br>Heterogeneity: Chi <sup>2</sup> =<br>Test for overall effect:<br>Study or Subgroup<br>Bao H 2019<br>Chen C 2018<br>Chen V 2018<br>Chen V 2018   | 588<br>19.51, df<br>Z = 6.91<br>Kangla<br>Events<br>25<br>16<br>17<br>22  | = 31 (P<br>(P < 0.0<br>iite<br><u>Total</u><br>31<br>30<br>44<br>51   | 415<br>= 0.95);   <sup>2</sup><br>60001)<br>Chemothe<br>Events<br>17<br>11<br>15<br>18   | erapy<br><u>Total</u><br>31<br>30<br>44<br>51   | Weight<br>2.4%<br>2.3%<br>3.4%<br>4.0%   | Risk Difference<br>M-H, Fixed, 95% CI<br>0.26 (0.03, 0.48)<br>0.17 (-0.08, 0.41)<br>0.05 (-0.16, 0.25)<br>0.08 (-0.11, 0.27)   | 0.01 0.1 10 100<br>Favours (Kanglaite) Favours (Chemotherapy)<br>Risk Difference<br>M-H, Fixed, 95% Cl  |
| Total events<br>Heterogeneity: Chi <sup>2</sup> =<br>Test for overall effect:<br>Study or Subgroup<br>Bao H 2019<br>Chen C 2018<br>Chen W 2018<br>Chen Y 2018<br>Chen Y 2013  | 588<br>19.51, df<br>Z = 6.91 (<br>Kangla<br>Events<br>25<br>16<br>17<br>22<br>13  | = 31 (P<br>P < 0.0<br>iite<br><u>Total</u><br>30<br>44<br>51<br>28  | 415<br>= 0.95);   <sup>2</sup><br>60001)<br>Chemothe<br>Events<br>17<br>11<br>15<br>18<br>9  | erapy<br><u>Total</u><br>31<br>30<br>44<br>51<br>27   | Weight<br>2.4%<br>2.3%<br>3.4%<br>4.0%<br>2.1%   | Risk Difference<br>M-H, Fixed, 95% CI<br>0.26 (0.03, 0.48)<br>0.17 (-0.08, 0.41)<br>0.05 (-0.16, 0.25)<br>0.08 (-0.11, 0.27)<br>0.13 (-0.13, 0.39)   | 0.01 0.1 10 100<br>Favours (Kanglaite) Favours (Chemotherapy)<br>Risk Difference<br>M-H, Fixed, 95% Cl  |
| Total events<br>Heterogeneity: Chi <sup>2</sup> =<br>Test for overall effect:<br>Study or Subgroup<br>Bao H 2019<br>Chen C 2018<br>Chen V 2018<br>Chen V 2018<br>Chen Y 2003<br>Chen YZ 2003<br>Guan XO 2009  | 588<br>19.51, df<br>Z = 6.91 (<br>Kangla<br>Events<br>25<br>16<br>17<br>22<br>13<br>5   | = 31 (P<br>(P < 0.0<br>iite<br><u>Total</u><br>31<br>30<br>44<br>51<br>28<br>12   | 415<br>= 0.95);   <sup>2</sup><br>00001)<br>Chemothe<br>Events<br>17<br>11<br>15<br>18<br>9<br>4   | erapy<br><u>Total</u><br>31<br>30<br>44<br>51<br>27<br>12   | Weight<br>2.4%<br>2.3%<br>3.4%<br>4.0%<br>2.1%<br>0.9%   | Risk Difference<br>M-H, Fixed, 95% CI<br>0.28 (0.03, 0.48)<br>0.17 (-0.08, 0.41)<br>0.05 (-0.16, 0.25)<br>0.08 (-0.11, 0.27)<br>0.13 (-0.13, 0.39)<br>0.08 (-0.30, 0.47)   | 0.01 0.1 10 100<br>Favours (Kanglaite) Favours (Chemotherapy)<br>Risk Difference<br>M-H, Fixed, 95% CI  |
| Total events<br>Heterogeneity: Chi <sup>a</sup> =<br>Test for overall effect:<br>Study or Subgroup<br>Bao H 2019<br>Chen C 2018<br>Chen V 2018<br>Chen YZ 2003<br>Guan XO 2009<br>He LT 2017  | 588<br>19.51, df<br>Z = 6.91 (<br>Kangla<br>Events<br>25<br>16<br>17<br>22<br>13<br>5<br>22   | = 31 (P<br>P < 0.0<br>inite<br>Total<br>31<br>30<br>44<br>51<br>28<br>12<br>54  | 415<br>= 0.95);   <sup>2</sup><br>00001)<br>Chemothe<br>Events<br>17<br>11<br>15<br>18<br>9<br>4<br>11   | = 0%<br>erapy<br><u>Total</u><br>31<br>30<br>44<br>51<br>27<br>12<br>54   | Weight<br>2.3%<br>3.4%<br>4.0%<br>2.1%<br>0.9%<br>4.2%   | Risk Difference<br>M-H, Fixed, 95% CI<br>0.26 (0.03, 0.48)<br>0.05 (-0.16, 0.25)<br>0.08 (-0.11, 0.27)<br>0.13 (-0.13, 0.39)<br>0.08 (-0.30, 0.47)<br>0.20 (0.03, 0.37)  | 0.01 0.1 10 100<br>Favours (Kanglaite) Favours (Chemotherapy)<br>Risk Difference<br>M-H, Fixed, 95% Cl  |
| Total events<br>Heterogeneity: Chi <sup>2</sup> =<br>Test for overall effect:<br>Study or Subgroup<br>Bao H 2019<br>Chen C 2018<br>Chen V 2018<br>Chen V 2018<br>Chen Y 2003<br>Guan XO 2009<br>He LT 2017<br>Huang ZB 2010   | 588<br>19.51, df<br>Z = 6.91 (<br>Kangla<br><u>Events</u><br>25<br>16<br>17<br>22<br>13<br>5<br>22<br>19  | = 31 (P<br>P < 0.0<br>iite<br><u>Total</u><br>31<br>30<br>44<br>51<br>28<br>12<br>54<br>35  | 415<br>= 0.95);   <sup>2</sup><br>00001)<br>Chemothe<br>Events<br>17<br>11<br>15<br>18<br>9<br>4<br>11<br>13   | = 0%<br>erapy<br><u>Total</u><br>31<br>30<br>44<br>51<br>27<br>12<br>54<br>35   | Weight<br>2.4%<br>2.3%<br>4.0%<br>2.1%<br>0.9%<br>4.2%<br>2.7%   | Risk Difference<br>M-H, Fixed, 95% CI<br>0.26 (0.03, 0.48)<br>0.17 (-0.08, 0.41)<br>0.05 (-0.16, 0.25)<br>0.08 (-0.13, 0.25)<br>0.08 (-0.13, 0.39)<br>0.08 (-0.30, 0.47)<br>0.20 (0.03, 0.37)<br>0.20 (0.03, 0.40)   | 0.01 0.1 10 100<br>Favours (Kanglaite) Favours (Chemotherapy)<br>Risk Difference<br>M.H. Fixed, 95% Cl  |
| Total events<br>Heterogeneity: Chi <sup>2</sup> =<br>Test for overall effect:<br>Study or Subgroup<br>Bao H 2019<br>Chen C 2018<br>Chen V 2018<br>Chen V 2003<br>Guan XO 2009<br>He LT 2017<br>Huang ZB 2010<br>Jia JN 2018   | 588<br>19.51, df<br>Z = 6.91 (<br>Kangla<br><u>Events</u><br>25<br>16<br>17<br>22<br>13<br>5<br>22<br>13<br>5<br>22<br>18   | = 31 (P<br>P < 0.0<br>inite<br>Total<br>31<br>30<br>44<br>51<br>28<br>12<br>54<br>35<br>31  | 415<br>= 0.95); P<br>00001)<br>Chemoth<br>Events<br>17<br>11<br>15<br>18<br>9<br>4<br>11<br>13<br>10   | erapy<br>Total<br>31<br>30<br>44<br>51<br>27<br>12<br>54<br>35<br>31  | Weight<br>2.4%<br>2.3%<br>3.4%<br>4.0%<br>2.1%<br>0.9%<br>4.2%<br>2.7%<br>2.4%   | Risk Difference<br><u>M-H, Fixed, 95% CI</u><br>0.28 (0.03, 0.48)<br>0.17 (-0.08, 0.41)<br>0.05 (-0.16, 0.25)<br>0.08 (-0.11, 0.27)<br>0.13 (-0.13, 0.39)<br>0.08 (-0.30, 0.47)<br>0.20 (0.03, 0.37)<br>0.17 (-0.06, 0.40)<br>0.16 (-0.08, 0.40)   | 0.01 0.1 10 100<br>Favours [Kanglaite] Favours [Chemotherapy]<br>Risk Difference<br>M-H, Fixed, 95% Cl  |
| Total events<br>Heterogeneity: Chi <sup>2</sup> =<br>Test for overall effect:<br>Study or Subgroup<br>Bao H 2019<br>Chen C 2018<br>Chen V 2018<br>Chen V 2018<br>Chen Y 2018<br>Chen Y 2003<br>Guan X0 2009<br>He LT 2017<br>Huang ZB 2010<br>Jia JN 2018<br>Li HY 2017   | 588<br>19.51, df<br>Z = 6.91 (<br>Kangla<br>Pvents<br>25<br>16<br>17<br>22<br>13<br>5<br>22<br>13<br>5<br>22<br>18<br>15<br>18<br>15<br>16  | = 31 (P<br>P < 0.0<br>iite<br><u>Total</u><br>31<br>30<br>44<br>51<br>28<br>12<br>54<br>35<br>31<br>41  | 415<br>= 0.95); P<br>60001)<br>Chemoth<br>Events<br>17<br>11<br>15<br>18<br>9<br>4<br>11<br>13<br>10<br>14   | erapy<br>Total<br>31<br>30<br>44<br>51<br>27<br>12<br>54<br>35<br>31<br>41  | Weight<br>2.4%<br>2.3%<br>3.4%<br>4.0%<br>2.1%<br>0.9%<br>4.2%<br>2.4%<br>2.4%<br>3.2%   | Risk Difference<br>M-H, Fixed, 95% CI<br>0.28 (0.03, 0.48)<br>0.17 (-0.08, 0.41)<br>0.05 (-0.16, 0.25)<br>0.08 (-0.11, 0.27)<br>0.13 (-0.13, 0.39)<br>0.08 (-0.30, 0.47)<br>0.20 (0.03, 0.37)<br>0.17 (-0.06, 0.40)<br>0.16 (-0.08, 0.40)<br>0.05 (-0.16, 0.28)  | 0.01 0.1 10 100<br>Favours [Kanglaite] Favours [Chemotherapy]<br>Risk Difference<br>M-H, Fixed, 95% Cl  |
| Total events<br>Heterogeneity: Chi <sup>2</sup> =<br>Test for overall effect:<br>Study or Subgroup<br>Bao H 2019<br>Chen C 2018<br>Chen V 2018<br>Chen V 2018<br>Chen YZ 2003<br>Guan XO 2009<br>He LT 2017<br>Huang ZB 2010<br>Jia JN 2018<br>LI HY 2017<br>LI L2012   | 588<br>19.51, df<br>Z = 6.91 (<br>Kangla<br>Events<br>25<br>16<br>17<br>22<br>13<br>5<br>22<br>19<br>15<br>16<br>8<br>8   | = 31 (P<br>P < 0.0<br>iite<br><u>Total</u><br>31<br>30<br>44<br>51<br>28<br>12<br>54<br>35<br>31<br>41<br>38  | 415<br>= 0.95); F<br>0001)<br>Chemoth<br>Events<br>17<br>11<br>15<br>18<br>9<br>4<br>11<br>13<br>10<br>14<br>9   | erapy<br>Total<br>31<br>30<br>44<br>51<br>27<br>12<br>54<br>35<br>35<br>31<br>41<br>40  | Weiaht<br>2.4%<br>2.3%<br>3.4%<br>2.1%<br>0.9%<br>4.2%<br>2.7%<br>2.4%<br>3.2%<br>3.0%   | Risk Difference<br>M-H, Fixed, 95% CI<br>0.26 (0.03, 0.48)<br>0.05 (-0.16, 0.25)<br>0.08 (-0.11, 0.27)<br>0.13 (-0.13, 0.39)<br>0.08 (-0.30, 0.47)<br>0.20 (-0.03, 0.37)<br>0.17 (-0.06, 0.40)<br>0.16 (-0.08, 0.40)<br>0.05 (-0.16, 0.28)<br>-0.01 (-0.20, 0.17)  | 0.01 0.1 10 100<br>Favours (Kanglaite) Favours (Chemotherapy)<br>Risk Difference<br>M-H, Fixed, 95% Cl  |
| Total events<br>Heterogeneity: Chi <sup>2</sup> =<br>Test for overall effect:<br>Study or Subgroup<br>Bao H 2019<br>Chen C 2018<br>Chen V 2018<br>Chen V 2018<br>Chen V 2003<br>Guan XO 2009<br>He LT 2017<br>Huang ZB 2010<br>Jia JN 2018<br>Li HV 2017<br>Li L 2012<br>Li XD 2016   | 588<br>19.51, df<br>Z = 6.91 (<br>Kangla<br><u>Events</u><br>25<br>166<br>17<br>22<br>13<br>5<br>22<br>22<br>13<br>5<br>22<br>22<br>18<br>15<br>16<br>8<br>8<br>24  | = 31 (P<br>P < 0.0<br>iite<br><u>Total</u><br>31<br>30<br>44<br>51<br>28<br>12<br>54<br>35<br>31<br>38<br>39  | 415<br>= 0.95), i <sup>2</sup><br>0001)<br>Chemoth<br>Events<br>17<br>11<br>15<br>18<br>9<br>4<br>11<br>13<br>10<br>14<br>9<br>14  | erapy<br>Total<br>31<br>30<br>44<br>51<br>27<br>12<br>54<br>35<br>31<br>41<br>40<br>39  | Weiaht<br>2.4%<br>2.3%<br>3.4%<br>2.1%<br>0.9%<br>4.2%<br>2.7%<br>2.4%<br>3.2%<br>3.0%<br>3.0%   | Risk Difference<br>M-H, Fixed, 95% CI<br>0.26 (0.03, 0.48)<br>0.17 (-0.08, 0.41)<br>0.05 (-0.16, 0.25)<br>0.08 (-0.13, 0.39)<br>0.08 (-0.30, 0.47)<br>0.20 (0.03, 0.37)<br>0.20 (0.03, 0.37)<br>0.17 (-0.05, 0.40)<br>0.16 (-0.08, 0.40)<br>0.05 (-0.16, 0.28)<br>0.01 (-0.20, 0.17)<br>0.26 (0.04, 0.47)  | 0.01 0.1 10 100 Favours (Kanglaite) Favours (Chemotherapy) Risk Difference M.H. Fixed, 95% Cl           |
| Total events<br>Heterogeneity: Chi <sup>2</sup> =<br>Test for overall effect:<br>Study or Subgroup<br>Bao H 2019<br>Chen C 2018<br>Chen V 2018<br>Chen V 2003<br>Guan XO 2009<br>He LT 2017<br>Huang ZB 2010<br>Jia JN 2018<br>Li HY 2017<br>Li L2012<br>Li XD 2016<br>Liang J 2018   | 588<br>19.51, df<br>Z = 6.91 (<br>Kangla<br><u>Events</u><br>25<br>16<br>17<br>22<br>13<br>5<br>22<br>13<br>5<br>22<br>18<br>15<br>16<br>15<br>16<br>15<br>16<br>24<br>23<br>24<br>23   | = 31 (P<br>P < 0.0<br>ite<br>Total<br>30<br>44<br>51<br>28<br>12<br>54<br>35<br>31<br>41<br>38<br>39<br>40  | 415<br>'= 0.95); /*<br>00001)<br>Chemothe<br><u>Events</u><br>17<br>11<br>15<br>18<br>9<br>4<br>11<br>13<br>10<br>14<br>9<br>4<br>10<br>14   | erapy<br><u>Total</u><br>31<br>30<br>44<br>51<br>27<br>12<br>54<br>35<br>31<br>41<br>40<br>39<br>40   | Weight<br>2.4%<br>2.3%<br>3.4%<br>4.0%<br>2.1%<br>0.9%<br>4.2%<br>2.7%<br>2.4%<br>3.2%<br>3.0%<br>3.0%<br>3.1%   | Risk Difference<br>M-H, Fixed, 95% CI<br>0.28 (0.03, 0.48)<br>0.17 (+0.08, 0.41)<br>0.05 (+0.16, 0.25)<br>0.08 (+0.11, 0.27)<br>0.13 (+0.13, 0.39)<br>0.08 (+0.30, 0.47)<br>0.20 (0.03, 0.37)<br>0.17 (+0.08, 0.40)<br>0.16 (+0.08, 0.40)<br>0.05 (+0.16, 0.26)<br>-0.01 (+0.20, 0.17)<br>0.26 [0.04, 0.47]<br>0.22 (0.12, 0.53)   | 0.01 0.1 10 100 Favours [Kanglaite] Favours [Chemotherapy] Risk Difference M-H, Fixed, 95% CI           |
| Total events<br>Heterogeneity: Chi <sup>P</sup> =<br>Test for overall effect:<br>Study or Subgroup<br>Bao H 2019<br>Chen C 2018<br>Chen V 2018<br>Chen V 2018<br>Chen V 2018<br>Chen V 2018<br>Guan XO 2009<br>He LT 2017<br>Huang ZB 2010<br>Jia JN 2018<br>Li HY 2017<br>Li L 2012<br>Li AD 2016<br>Liang JG 2014   | 588<br>19.51, df<br>Z = 6.91 (<br>Kangle<br><u>Events</u><br>225<br>16<br>17<br>22<br>18<br>15<br>22<br>18<br>15<br>16<br>16<br>8<br>22<br>23<br>18<br>15<br>16<br>16<br>18<br>19<br>24<br>24<br>23<br>21<br>0  | = 31 (P<br>P < 0.0<br>ite<br>Total<br>30<br>44<br>51<br>28<br>12<br>54<br>35<br>31<br>41<br>38<br>39<br>40<br>23  | 415<br>'= 0.95), i <sup>*</sup><br>00001)<br>Chemoth-<br>Events<br>17<br>11<br>15<br>18<br>9<br>4<br>11<br>13<br>10<br>14<br>9<br>14<br>9<br>14<br>9<br>14<br>8  | erapy<br>Total<br>31<br>30<br>44<br>51<br>27<br>12<br>54<br>35<br>31<br>41<br>40<br>39<br>40<br>20  | Weiaht<br>2.4%<br>2.3%<br>3.4%<br>4.0%<br>2.1%<br>4.2%<br>2.7%<br>2.4%<br>3.2%<br>3.0%<br>3.0%<br>3.1%<br>1.7%   | Risk Difference<br>M-H, Fixed, 95% CI<br>0.28 (0.03, 0.48)<br>0.17 (-0.08, 0.41)<br>0.05 (-0.16, 0.25)<br>0.08 (-0.11, 0.27)<br>0.13 (-0.13, 0.39)<br>0.08 (-0.03, 0.37)<br>0.17 (-0.06, 0.40)<br>0.16 (-0.08, 0.40)<br>0.05 (-0.16, 0.28)<br>-0.01 (-0.20, 0.17)<br>0.26 [0.04, 0.47]<br>0.26 [0.12, 0.53]<br>0.03 (-0.26, 0.33]  | 0.01 0.1 10 100 Favours [Kanglaite] Favours [Chemotherapy] Risk Difference M.H. Fixed, 95% Cl           |
| Total events<br>Heterogeneity: Chi <sup>2</sup> =<br>Test for overall effect:<br>Study or Subgroup<br>Bao H 2019<br>Chen C 2018<br>Chen V 2018<br>Chen V 2018<br>Chen YZ 2003<br>Guan X0 2009<br>He LT 2017<br>Huang ZB 2010<br>Jia JN 2018<br>Li HY 2017<br>Li L2012<br>Li XD 2016<br>Liang J 2018<br>Liang SG 2014<br>Liu F 2019  | 588<br>19.51, df<br>Z = 6.91 (<br>Kangle<br><u>Events</u><br>255<br>166<br>17<br>22<br>13<br>5<br>22<br>18<br>15<br>16<br>16<br>8<br>24<br>23<br>10<br>0<br>40  | = 31 (P<br>P < 0.0<br>inte<br><u>Total</u><br>31<br>30<br>44<br>51<br>28<br>12<br>54<br>35<br>31<br>41<br>38<br>39<br>40<br>23<br>63  | 415<br>(= 0.95); (=<br>0001)<br>Chemothin<br>Events<br>17<br>11<br>15<br>5<br>18<br>9<br>4<br>11<br>13<br>10<br>14<br>13<br>10<br>14<br>10<br>14<br>10<br>14<br>10<br>14<br>26   | = 0%<br>Total<br>31<br>30<br>44<br>51<br>27<br>12<br>54<br>35<br>31<br>41<br>40<br>39<br>40<br>20<br>63   | Weialitt<br>2.4%<br>2.3%<br>3.4%<br>2.1%<br>2.1%<br>2.4%<br>2.7%<br>2.4%<br>3.2%<br>3.0%<br>3.0%<br>3.0%<br>3.0%<br>4.9%   | Risk Difference<br>M-H, Fixed, 95% CI<br>0.28 (0.03, 0.48)<br>0.17 (-0.08, 0.41)<br>0.05 (-0.14), 0.25]<br>0.08 (-0.11, 0.27)<br>0.13 (-0.13, 0.38)<br>0.08 (-0.30, 0.47)<br>0.20 (0.03, 0.37)<br>0.17 (-0.06, 0.40)<br>0.16 (-0.08, 0.40)<br>0.16 (-0.08, 0.40)<br>0.16 (-0.16, 0.26)<br>-0.01 (-0.20, 0.17)<br>0.26 [0.04, 0.47]<br>0.26 [0.04, 0.47]<br>0.26 [0.04, 0.47]<br>0.22 (0.12, 0.53)<br>0.03 (-0.26, 0.33)<br>0.22 (0.05, 0.39]   | 0.01 0.1 10 100 Favours [Kanglaite] Favours [Chemotherapy] Risk Difference M.H. Fixed, 95% Cl           |
| Total events<br>Heterogeneity: Chi <sup>2</sup> =<br>Test for overall effect:<br>Bao H 2019<br>Chen C 2018<br>Chen V 2018<br>Chen V 2018<br>Chen V 2003<br>Guan XO 2009<br>He LT 2017<br>Huang ZB 2010<br>Jia JN 2018<br>Li HV 2017<br>Li L 2012<br>Li XD 2016<br>Liang J 2018<br>Liang SG 2014<br>Liu F 2019<br>Liu JQ 2011  | 588<br>19.51, df<br>Z = 6.91 (<br>Kangla<br>Events<br>225<br>16<br>17<br>222<br>13<br>5<br>22<br>13<br>5<br>22<br>13<br>15<br>15<br>16<br>8<br>8<br>24<br>23<br>10<br>40<br>0<br>15   | = 31 (P < 0.0<br>inite<br>Total<br>31<br>30<br>44<br>51<br>52<br>82<br>54<br>35<br>31<br>31<br>38<br>39<br>40<br>23<br>83<br>83<br>83<br>83<br>83<br>83<br>83   | 415<br>(= 0.95);  *<br>60001)<br>Chemoth<br>Events<br>17<br>11<br>15<br>18<br>9<br>4<br>13<br>10<br>14<br>9<br>4<br>13<br>10<br>14<br>9<br>14<br>10<br>8<br>26<br>12   | erapy<br>Total<br>31<br>30<br>44<br>51<br>27<br>12<br>54<br>31<br>41<br>40<br>39<br>40<br>20<br>63<br>35  | Weiaht<br>2.4%<br>2.3%<br>3.4%<br>2.1%<br>0.9%<br>4.2%<br>3.0%<br>3.0%<br>3.0%<br>3.1%<br>3.1%<br>4.8%<br>2.7%   | Risk Difference<br>M-H, Fixed, 95% CI<br>0.26 (0.03, 0.48)<br>0.17 (-0.08, 0.41)<br>0.05 (-0.16, 0.25)<br>0.08 (-0.11, 0.27)<br>0.13 (-0.13, 0.39)<br>0.08 (-0.30, 0.37)<br>0.17 (-0.06, 0.40)<br>0.16 (-0.08, 0.40)<br>0.05 (-0.16, 0.20, 0.17)<br>0.26 (0.04, 0.47)<br>0.32 (0.12, 0.53)<br>0.32 (0.12, 0.53)<br>0.32 (0.25, 0.39)<br>0.29 (-0.5, 0.39)<br>0.29 (-0.14, 0.31)  | 0.01 0.1 10 100 Favours (Kanglaite) Favours (Chemotherapy) Risk Difference M-H, Fixed, 95% Cl           |
| Total events<br>Heterogeneity: Chi <sup>2</sup> =<br>Test for overall effect:<br>Bao H 2019<br>Chen C 2018<br>Chen V 2018<br>Chen V 2018<br>Chen V 2003<br>Guan XO 2009<br>He LT 2017<br>Huang ZB 2010<br>Jia JN 2018<br>Li HY 2017<br>Li L 2012<br>Li XD 2016<br>Liang J 2018<br>Liang SG 2014<br>Liu JQ 2011<br>Liu XD 2015   | 588<br>19.51, df<br>Z = 6.91 (<br><u>Rends</u><br>255<br>16<br>17<br>22<br>18<br>16<br>17<br>22<br>13<br>5<br>22<br>13<br>5<br>22<br>13<br>15<br>15<br>16<br>16<br>8<br>24<br>40<br>23<br>10<br>40<br>40<br>40<br>27  | = 31 (P < 0.0<br>inite<br>Total<br>31<br>30<br>44<br>51<br>52<br>54<br>31<br>32<br>54<br>35<br>31<br>38<br>39<br>940<br>23<br>39<br>940<br>23<br>35<br>55   | 415<br>= 0.95); /*<br>0001)<br>Chemothin<br>Events<br>17<br>11<br>15<br>15<br>18<br>9<br>4<br>4<br>11<br>13<br>10<br>14<br>11<br>13<br>10<br>14<br>14<br>10<br>8<br>26<br>22<br>17   | = 0%<br>Total<br>31<br>30<br>44<br>51<br>27<br>12<br>54<br>35<br>31<br>41<br>40<br>39<br>40<br>20<br>63<br>35<br>55   | Weialtt<br>2.4%<br>2.3%<br>3.4%<br>4.0%<br>2.1%<br>0.9%<br>4.2%<br>2.7%<br>3.2%<br>3.0%<br>3.0%<br>3.1%<br>1.7%<br>4.9%<br>2.7%<br>4.3%  | Risk Difference<br>M-H, Fixed, 95% CI<br>0.28 (0.03, 0.48)<br>0.17 (+0.08, 0.41)<br>0.05 (+0.16, 0.25)<br>0.08 (+0.11, 0.27)<br>0.13 (+0.13, 0.39)<br>0.08 (+0.30, 0.47)<br>0.20 (0.03, 0.37)<br>0.17 (+0.08, 0.40)<br>0.16 (+0.08, 0.40)<br>0.16 (+0.08, 0.40)<br>0.05 (+0.16, 0.26)<br>-0.01 (+0.20, 0.17)<br>0.26 (0.04, 0.47)<br>0.32 (0.12, 0.53)<br>0.03 (+0.26, 0.33)<br>0.22 (0.05, 0.38)<br>0.09 (+0.14, 0.31)<br>0.18 (0.00, 0.36)   | 0.01 0.1 10 100 Favours (Kanglaite) Favours (Chemotherapy) Risk Difference M-H, Fixed, 95% CI           |
| Total events<br>Heterogeneity: Chi <sup>2</sup> =<br>Test for overall effect:<br>Study or Subgroup<br>Bao H 2019<br>Chen C 2018<br>Chen V 2018<br>Chen Y 2018<br>Chen YZ 2003<br>Guan XO 2009<br>He LT 2017<br>Huang ZB 2010<br>Jia JN 2018<br>Li HY 2017<br>Li L 2012<br>Li AD 2016<br>Liang J 2018<br>Liang SG 2014<br>Liu JQ 2011<br>Liu XH 2016<br>Liu Y 2015   | 588<br>19.51, df<br>Z = 6.91 (<br>Kangla<br>Events<br>25<br>16<br>17<br>22<br>13<br>5<br>22<br>22<br>13<br>5<br>22<br>22<br>13<br>5<br>22<br>22<br>13<br>15<br>16<br>15<br>16<br>15<br>16<br>15<br>15<br>16<br>15<br>15<br>16<br>15<br>15<br>16<br>15<br>17<br>27<br>18<br>15<br>16<br>16<br>17<br>17<br>18<br>18<br>19<br>19<br>19<br>19<br>19<br>19<br>19<br>19<br>19<br>19<br>19<br>19<br>19   | = 31 (P < 0.0<br>inite<br>Total<br>31<br>30<br>44<br>51<br>54<br>54<br>35<br>54<br>31<br>38<br>39<br>40<br>23<br>83<br>39<br>40<br>23<br>83<br>55<br>55   | 415<br>= 0.95); /*<br>00001)<br>Chemothin<br>Events<br>17<br>11<br>15<br>5<br>18<br>9<br>4<br>4<br>11<br>13<br>10<br>14<br>11<br>13<br>10<br>14<br>10<br>14<br>10<br>14<br>10<br>14<br>10<br>14  | = 0%.<br>Total<br>31<br>30<br>44<br>51<br>27<br>12<br>54<br>31<br>41<br>40<br>39<br>940<br>20<br>63<br>355<br>43  | Weight<br>2.4%<br>2.3%<br>3.4%<br>2.1%<br>0.9%<br>2.7%<br>2.4%<br>3.0%<br>3.0%<br>3.0%<br>3.0%<br>3.0%<br>3.0%<br>3.0%<br>3.0  | Risk Difference<br>M-H, Fixed, 95% CI<br>0.28 (0.03, 0.48)<br>0.17 (-0.08, 0.41)<br>0.05 (-0.16, 0.25)<br>0.08 (-0.11, 0.27)<br>0.13 (-0.13, 0.39)<br>0.08 (-0.30, 0.47)<br>0.20 (0.03, 0.37)<br>0.17 (-0.06, 0.40)<br>0.16 (-0.08, 0.40)<br>0.05 (-0.16, 0.28)<br>-0.01 (-0.20, 0.17)<br>0.26 [0.04, 0.47]<br>0.26 [0.04, 0.47]<br>0.26 [0.04, 0.47]<br>0.26 [0.05, 0.38]<br>0.03 (-0.26, 0.33]<br>0.22 (0.05, 0.38]<br>0.09 (-0.14, 0.31]<br>0.18 [0.00, 0.36]<br>0.08 (-0.11, 0.30]   | 0.01 0.1 10 100 Favours [Kanglaite] Favours [Chemotherapy] Risk Difference M.H. Fixed, 95% Cl           |
| Total events<br>Heterogeneity: Chi <sup>2</sup> =<br>Test for overall effect:<br>Study or Subgroup<br>Bao H 2019<br>Chen C 2018<br>Chen V 2018<br>Chen V 2018<br>Chen YZ 2003<br>Guan X0 2009<br>He LT 2017<br>Huang ZB 2010<br>Jia JN 2018<br>Li HY 2017<br>Li L 2012<br>Li XD 2018<br>Liang SG 2014<br>Liung 2015<br>Liung 2015<br>Liung 2015<br>Liong SG 2017  | 588<br>19.51, df<br>Z = 6.91 (<br><b>Kangla</b><br><b>Events</b><br>25<br>16<br>17<br>22<br>13<br>5<br>22<br>13<br>13<br>5<br>22<br>18<br>15<br>16<br>8<br>24<br>23<br>10<br>10<br>10<br>40<br>40<br>15<br>27<br>18   | = 31 (P < 0.0<br>inite<br>Total<br>31<br>30<br>44<br>51<br>28<br>54<br>54<br>54<br>35<br>31<br>38<br>39<br>99<br>40<br>23<br>35<br>55<br>54<br>3<br>42<br>23<br>55<br>54<br>34<br>24<br>23  | 415<br>(= 0.95); (=<br>0001)<br>Chemoth<br>Events<br>17<br>11<br>15<br>5<br>18<br>9<br>4<br>11<br>13<br>10<br>14<br>9<br>4<br>11<br>13<br>10<br>14<br>9<br>4<br>11<br>13<br>10<br>14<br>9<br>26<br>12<br>17<br>14<br>15<br>15  | = 0%<br><b>Total</b><br>31<br>30<br>44<br>51<br>27<br>12<br>54<br>35<br>54<br>41<br>40<br>39<br>40<br>20<br>63<br>35<br>55<br>54<br>34<br>40<br>40<br>40<br>40<br>40<br>40<br>40<br>40<br>40<br>4   | Weight<br>2.4%<br>2.3%<br>3.4%<br>2.1%<br>0.9%<br>4.2%<br>2.7%<br>2.4%<br>3.0%<br>3.0%<br>3.0%<br>3.0%<br>3.0%<br>3.0%<br>3.0%<br>3.0  | Risk Difference<br>M-H, Fixed, 95% CI<br>0.28 (0.03, 0.48)<br>0.17 (+0.08, 0.41)<br>0.05 (-0.16, 0.25)<br>0.08 (-0.11, 0.27)<br>0.13 (-0.13, 0.38)<br>0.08 (-0.30, 0.47)<br>0.20 (0.03, 0.37)<br>0.17 (+0.06, 0.40)<br>0.16 (+0.08, 0.40)<br>0.16 (+0.08, 0.40)<br>0.05 (-0.16, 0.26)<br>-0.01 (+0.20, 0.17)<br>0.25 (0.04, 0.47)<br>0.25 (0.04, 0.47)<br>0.25 (0.04, 0.43)<br>0.03 (+0.26, 0.33)<br>0.22 (0.05, 0.39)<br>0.09 (+0.14, 0.31)<br>0.08 (+0.14, 0.30)<br>0.03 (+0.18, 0.24)   | 0.01 0.1 10 100 Favours [Kanglaite] Favours [Chemotherapy] Risk Difference M.H. Fixed, 95% Cl           |
| Total events<br>Heterogeneity: Chi <sup>2</sup> =<br>Test for overall effect:<br>Bao H 2019<br>Chen C 2018<br>Chen V 2018<br>Chen V 2018<br>Chen V 2003<br>Guan XO 2009<br>He LT 2017<br>Huang ZB 2010<br>Jia JN 2018<br>Li HV 2017<br>Li L 2012<br>Li XD 2016<br>Liang SG 2014<br>Liu F 2019<br>Liu JQ 2011<br>Liu XH 2015<br>Liong SG 2017<br>Mu Q 2018   | 588<br>19.51, df<br>Z = 6.91 (<br>Kangla<br>Events<br>25<br>16<br>17<br>22<br>13<br>5<br>22<br>13<br>5<br>22<br>13<br>15<br>16<br>15<br>16<br>8<br>8<br>24<br>23<br>10<br>40<br>40<br>15<br>27<br>18<br>17<br>27<br>27<br>24  | = 31 (P < 0.0<br>inte<br>Total<br>31<br>30<br>44<br>51<br>12<br>54<br>35<br>54<br>31<br>32<br>54<br>33<br>39<br>940<br>23<br>39<br>940<br>23<br>35<br>55<br>55<br>55<br>43<br>42<br>47<br>77  | 415<br>(= 0.95); /=<br>0001)<br>Chemoth<br>Events<br>17<br>11<br>15<br>18<br>9<br>4<br>13<br>10<br>14<br>9<br>4<br>13<br>10<br>14<br>13<br>10<br>14<br>10<br>8<br>26<br>12<br>17<br>14<br>15<br>12<br>12<br>17   | = 0%.<br>Total<br>31<br>30<br>44<br>51<br>27<br>12<br>27<br>12<br>35<br>31<br>41<br>40<br>39<br>40<br>40<br>35<br>55<br>43<br>35<br>55<br>43<br>40<br>40<br>40<br>40<br>40<br>40<br>40<br>40<br>40<br>40  | Weight<br>2.4%<br>3.4%<br>4.0%<br>2.1%<br>0.9%<br>4.2%<br>2.4%<br>3.0%<br>3.0%<br>3.0%<br>3.1%<br>4.8%<br>3.1%<br>4.3%<br>3.3%<br>3.3%<br>3.3%<br>3.8%   | Risk Difference<br><u>M-H, Fixed, 95% CI</u><br>0.28 (0.03, 0.48)<br>0.17 (-0.08, 0.41)<br>0.05 (-0.16, 0.25)<br>0.08 (-0.11, 0.27)<br>0.13 (-0.13, 0.39)<br>0.08 (-0.30, 0.47)<br>0.20 (0.03, 0.37)<br>0.17 (-0.06, 0.40)<br>0.16 (-0.08, 0.40)<br>0.05 (-0.16, 0.26)<br>-0.01 (-0.20, 0.17)<br>0.25 (0.04, 0.47)<br>0.22 (0.05, 0.40)<br>0.03 (-0.26, 0.33)<br>0.09 (-0.14, 0.31)<br>0.18 (0.00, 0.36)<br>0.03 (-0.18, 0.24)<br>0.03 (-0.18, 0.24)<br>0.03 (-0.18, 0.24)<br>0.21 (0.02, 0.41)  | 0.01 0.1 10 100 Favours (Kanglaite) Favours (Chemotherapy) Risk Difference M-H, Fixed, 95% Cl           |
| Total events<br>Heterogeneity: $Chi^2$ =<br>Test for overall effect:<br>Bao H 2019<br>Chen C 2018<br>Chen V 2018<br>Chen V 2018<br>Chen V 2003<br>Guan XO 2009<br>He LT 2017<br>Huang ZB 2010<br>Jia JN 2018<br>Li HY 2017<br>Li L 2012<br>Li XD 2018<br>Liang SG 2014<br>Liang SG 2014<br>Liu JQ 2011<br>Liu XH 2015<br>Long SG 2017<br>Mu Q 2018<br>Sun SQ 2012   | 588<br>19.51, df<br>Z = 6.91 (<br>Kangla<br>Events<br>25<br>16<br>17<br>22<br>13<br>5<br>22<br>18<br>15<br>16<br>15<br>16<br>15<br>16<br>15<br>16<br>15<br>22<br>18<br>15<br>16<br>15<br>16<br>15<br>16<br>15<br>16<br>15<br>16<br>15<br>16<br>17<br>22<br>18<br>15<br>16<br>17<br>22<br>18<br>16<br>17<br>22<br>18<br>16<br>17<br>22<br>18<br>17<br>22<br>18<br>17<br>22<br>18<br>17<br>22<br>18<br>17<br>22<br>18<br>17<br>22<br>18<br>17<br>22<br>18<br>17<br>22<br>18<br>17<br>22<br>18<br>17<br>22<br>18<br>17<br>22<br>18<br>18<br>17<br>22<br>18<br>18<br>19<br>17<br>22<br>18<br>18<br>19<br>18<br>19<br>18<br>19<br>18<br>19<br>19<br>19<br>19<br>19<br>19<br>19<br>19<br>19<br>19<br>19<br>19<br>19   | = 31 (P < 0.0<br>inte<br>Total<br>31<br>30<br>44<br>54<br>35<br>54<br>35<br>31<br>41<br>41<br>38<br>39<br>40<br>23<br>39<br>40<br>23<br>35<br>55<br>43<br>35<br>55<br>43<br>42<br>47<br>355   | 415<br>40.95); /*<br>0001)<br>Chemothin<br>Events<br>17<br>11<br>15<br>18<br>9<br>4<br>11<br>13<br>10<br>14<br>10<br>14<br>10<br>8<br>26<br>12<br>17<br>14<br>15<br>14<br>15<br>14<br>15<br>14<br>15<br>16<br>17<br>11<br>10<br>10<br>10<br>10<br>10<br>10<br>10<br>10<br>10   | = 0%.<br><b>Total</b><br>31<br>30<br>44<br>51<br>27<br>12<br>54<br>35<br>31<br>41<br>40<br>20<br>63<br>35<br>55<br>55<br>43<br>40<br>40<br>40<br>40<br>40<br>40<br>40<br>41<br>40<br>41<br>40<br>41<br>40<br>41<br>40<br>41<br>40<br>41<br>40<br>41<br>40<br>41<br>40<br>41<br>40<br>41<br>40<br>41<br>40<br>41<br>40<br>41<br>40<br>41<br>40<br>41<br>40<br>40<br>40<br>40<br>40<br>40<br>40<br>40<br>40<br>40 | Weight<br>2.4%<br>2.3%<br>3.4%<br>2.1%<br>0.9%<br>2.7%<br>2.4%<br>3.0%<br>3.0%<br>3.0%<br>3.1%<br>4.8%<br>2.7%<br>4.8%<br>3.3%<br>3.3%<br>3.3%<br>3.2%<br>3.3%                                 | Risk Difference<br>M-H, Fixed, 95% CI<br>0.28 (0.03, 0.48)<br>0.17 (-0.08, 0.41)<br>0.05 (-0.16, 0.25)<br>0.08 (-0.11, 0.27)<br>0.13 (-0.13, 0.39)<br>0.08 (-0.30, 0.47)<br>0.20 (0.03, 0.37)<br>0.17 (-0.06, 0.40)<br>0.16 (-0.08, 0.40)<br>0.05 (-0.16, 0.26)<br>-0.01 (-0.20, 0.17)<br>0.26 (0.04, 0.47)<br>0.22 (0.05, 0.39)<br>0.03 (-0.26, 0.33)<br>0.22 (0.05, 0.39)<br>0.09 (-0.14, 0.31)<br>0.18 (-0.00, 0.36)<br>0.09 (-0.14, 0.31)<br>0.18 (-0.00, 0.36)<br>0.09 (-0.14, 0.31)<br>0.3 (-0.18, 0.26)<br>0.09 (-0.14, 0.31)<br>0.3 (-0.18, 0.26)<br>0.09 (-0.14, 0.31)<br>0.3 (-0.18, 0.26)<br>0.09 (-0.14, 0.31)<br>0.3 (-0.18, 0.26)<br>0.3 (-0.2, 0.41)<br>0.11 (-0.11, 0.34)  | 0.01 0.1 10 100 Favours (Kanglaite) Favours (Chemotherapy) Risk Difference M.H. Fixed, 95% CI           |
| Total events<br>Heterogeneity: Chi <sup>2</sup> =<br>Test for overall effect:<br>Study or Subgroup<br>Bao H 2019<br>Chen C 2018<br>Chen V 2018<br>Chen V 2018<br>Chen Y 2003<br>Guan X0 2009<br>He LT 2017<br>Huang ZB 2010<br>Jia JN 2018<br>Li HY 2017<br>Li L 2012<br>Li XD 2016<br>Liang SG 2014<br>Liu F 2019<br>Liu JQ 2016<br>Liu Y 2015<br>Long SG 2017<br>Mu Q 2018<br>Sun SQ 2012<br>Wang CY 2018   | 588<br>19.51, df<br>Z= 6.91<br><b>Kangla</b><br><b>Events</b><br>25<br>16<br>17<br>22<br>13<br>5<br>22<br>13<br>5<br>22<br>18<br>15<br>16<br>16<br>16<br>16<br>8<br>8<br>24<br>23<br>10<br>10<br>15<br>24<br>23<br>10<br>10<br>15<br>24<br>15<br>16<br>16<br>15<br>16<br>16<br>17<br>24<br>18<br>16<br>16<br>17<br>24<br>18<br>16<br>16<br>17<br>24<br>18<br>16<br>16<br>17<br>24<br>18<br>16<br>19<br>17<br>24<br>25<br>18<br>16<br>16<br>17<br>27<br>27<br>27<br>27<br>27<br>27<br>27<br>27<br>27<br>27<br>27<br>27<br>27   | = 31 (P < 0.0<br>inte<br><u>Total</u><br>31<br>30<br>44<br>51<br>32<br>54<br>45<br>12<br>54<br>45<br>12<br>54<br>45<br>12<br>54<br>45<br>12<br>54<br>45<br>12<br>54<br>45<br>12<br>28<br>31<br>41<br>38<br>39<br>30<br>28<br>31<br>28<br>31<br>28<br>31<br>30<br>28<br>31<br>28<br>31<br>31<br>30<br>31<br>30<br>31<br>30<br>31<br>30<br>31<br>30<br>31<br>30<br>31<br>30<br>31<br>30<br>31<br>30<br>31<br>30<br>31<br>30<br>31<br>30<br>31<br>30<br>31<br>31<br>31<br>32<br>31<br>31<br>31<br>31<br>31<br>31<br>31<br>31<br>31<br>31   | 415<br>'= 0.95); /=<br>00001)<br>Chemothin<br>Events<br>17<br>11<br>15<br>5<br>18<br>9<br>4<br>4<br>11<br>13<br>10<br>14<br>13<br>10<br>14<br>10<br>14<br>10<br>8<br>26<br>127<br>17<br>14<br>15<br>14<br>11<br>17   | = 0%<br>trapy<br>Total<br>311<br>300<br>444<br>511<br>277<br>12<br>54<br>355<br>543<br>311<br>400<br>399<br>400<br>633<br>355<br>555<br>433<br>440<br>400<br>633<br>444<br>400<br>633<br>444<br>400<br>633<br>444<br>400<br>633<br>555<br>555<br>433<br>400<br>407<br>407<br>407<br>407<br>407<br>407<br>407  | Weight<br>2.4%<br>2.3%<br>3.4%<br>4.0%<br>2.1%<br>0.9%<br>2.2%<br>3.0%<br>3.0%<br>3.0%<br>3.0%<br>3.0%<br>3.0%<br>3.0%<br>3.0  | Risk Difference<br>M-H, Fixed, 95% CI<br>0.26 (0.03, 0.48)<br>0.17 (-0.08, 0.41)<br>0.56 (-0.16, 0.25)<br>0.08 (-0.11, 0.27)<br>0.13 (-0.13, 0.39)<br>0.08 (-0.30, 0.47)<br>0.20 (0.03, 0.37)<br>0.17 (-0.06, 0.40)<br>0.16 (-0.08, 0.40)<br>0.56 (-0.16, 0.28)<br>-0.01 (-0.20, 0.17)<br>0.26 [0.04, 0.47]<br>0.26 [0.04, 0.47]<br>0.26 [0.04, 0.47]<br>0.26 [0.04, 0.47]<br>0.22 (0.12, 0.53)<br>0.03 (-0.26, 0.33)<br>0.02 (-0.5, 0.33)<br>0.09 (-0.11, 0.30)<br>0.08 (-0.11, 0.30)<br>0.09 (-0.11, 0.30)<br>0.03 (-0.18, 0.24)<br>0.21 (-0.02, 0.41]<br>0.21 (-0.12, 0.41)<br>0.22 (-0.19, 0.23)   | D.01 0.1 10 100 Favours [Kanglaite] Favours [Chemotherapy] Risk Difference M.H. Fixed, 95% Cl           |
| Total events<br>Heterogeneity: Chi <sup>2</sup> =<br>Test for overall effect:<br>Study or Subgroup<br>Bao H 2019<br>Chen C 2018<br>Chen V 2018<br>Chen V 2018<br>Chen V 2018<br>Chen YZ 2003<br>Guan X0 2009<br>He LT 2017<br>Huang ZB 2010<br>Jia JN 2018<br>Li HY 2017<br>Li L 2012<br>Li XD 2018<br>Liang SG 2014<br>Liu F 2019<br>Liu JQ 2011<br>Liu Y 2015<br>Long SG 2017<br>Mu Q 2018<br>Sun SQ 2012<br>Wang CY 2018<br>Wang YZ 2018   | 588<br>19.51, df<br>Z = 6.91 (<br>Rangla<br>Events<br>25<br>16<br>17<br>22<br>13<br>5<br>22<br>13<br>5<br>22<br>13<br>5<br>22<br>13<br>15<br>16<br>8<br>8<br>24<br>23<br>10<br>40<br>40<br>40<br>15<br>27<br>18<br>17<br>27<br>18<br>10<br>15<br>16<br>16<br>16<br>17<br>22<br>13<br>13<br>5<br>22<br>13<br>15<br>16<br>17<br>22<br>13<br>13<br>5<br>15<br>16<br>17<br>22<br>13<br>13<br>15<br>16<br>17<br>17<br>22<br>13<br>13<br>15<br>16<br>17<br>22<br>13<br>13<br>15<br>16<br>17<br>22<br>13<br>13<br>15<br>16<br>17<br>22<br>13<br>13<br>15<br>16<br>17<br>17<br>22<br>13<br>13<br>15<br>16<br>17<br>17<br>22<br>13<br>13<br>15<br>16<br>17<br>17<br>22<br>13<br>13<br>15<br>16<br>16<br>17<br>17<br>22<br>13<br>13<br>15<br>16<br>16<br>17<br>17<br>22<br>13<br>13<br>15<br>16<br>16<br>17<br>17<br>22<br>13<br>13<br>15<br>16<br>16<br>17<br>17<br>22<br>13<br>13<br>15<br>16<br>16<br>17<br>17<br>22<br>13<br>13<br>15<br>16<br>16<br>17<br>17<br>22<br>13<br>13<br>15<br>16<br>16<br>17<br>17<br>22<br>13<br>13<br>15<br>16<br>16<br>16<br>17<br>17<br>22<br>13<br>13<br>15<br>16<br>16<br>17<br>17<br>22<br>13<br>13<br>15<br>16<br>16<br>17<br>17<br>22<br>18<br>18<br>15<br>16<br>16<br>17<br>17<br>22<br>19<br>18<br>15<br>16<br>18<br>17<br>17<br>22<br>18<br>18<br>15<br>16<br>16<br>16<br>16<br>17<br>17<br>17<br>17<br>18<br>18<br>16<br>16<br>16<br>17<br>17<br>18<br>18<br>16<br>16<br>17<br>17<br>18<br>18<br>16<br>16<br>17<br>17<br>18<br>18<br>18<br>19<br>18<br>19<br>19<br>18<br>19<br>19<br>18<br>19<br>19<br>19<br>19<br>19<br>19<br>19<br>19<br>19<br>19<br>19<br>19<br>19 | = 31 (P < 0.0<br>inte<br>Total<br>31<br>30<br>44<br>51<br>12<br>54<br>35<br>55<br>55<br>55<br>55<br>55<br>55<br>55<br>55<br>55  | 415<br>(= 0.95);  *<br>60001)<br>Chemothi<br>Events<br>17<br>11<br>15<br>18<br>9<br>4<br>11<br>13<br>10<br>14<br>13<br>10<br>14<br>13<br>10<br>14<br>10<br>14<br>10<br>14<br>10<br>14<br>11<br>13<br>10<br>14<br>11<br>13<br>10<br>14<br>11<br>13<br>10<br>14<br>11<br>13<br>10<br>14<br>11<br>13<br>10<br>14<br>11<br>13<br>10<br>14<br>11<br>13<br>14<br>11<br>13<br>14<br>14<br>14<br>15<br>15<br>17<br>17<br>17<br>17<br>17<br>17<br>17<br>17<br>17<br>17<br>17<br>17<br>17  | = 0%<br>trapy<br>Total<br>31<br>30<br>44<br>51<br>27<br>12<br>24<br>35<br>31<br>41<br>40<br>39<br>40<br>20<br>03<br>35<br>55<br>55<br>55<br>55<br>55<br>43<br>40<br>40<br>40<br>40<br>40<br>40<br>40<br>40<br>40<br>40  | Weight<br>2.4%<br>2.3%<br>3.4%<br>2.1%<br>0.9%<br>4.2%<br>2.7%<br>2.4%<br>3.0%<br>3.0%<br>3.0%<br>3.0%<br>3.1%<br>4.8%<br>2.7%<br>4.3%<br>3.2%<br>3.6%<br>3.2%<br>3.6%<br>3.3%<br>3.6%<br>3.3% | Risk Difference<br>M-H, Fixed, 95% CI<br>0.26 (0.03, 0.48)<br>0.17 (+0.08, 0.41)<br>0.05 (-0.16, 0.25)<br>0.08 (-0.11, 0.27)<br>0.13 (-0.13, 0.38)<br>0.08 (-0.30, 0.47)<br>0.20 (0.03, 0.37)<br>0.17 (+0.06, 0.40)<br>0.16 (-0.08, 0.40)<br>0.05 (-0.16, 0.28)<br>-0.01 (+0.20, 0.17)<br>0.26 (0.04, 0.47)<br>0.22 (0.05, 0.38)<br>0.03 (+0.26, 0.33)<br>0.03 (+0.26, 0.33)<br>0.09 (+0.14, 0.31)<br>0.08 (-0.11, 0.30)<br>0.03 (-0.18, 0.24)<br>0.21 (0.02, 0.41)<br>0.11 (-0.11, 0.34)<br>0.02 (-0.19, 0.23)<br>0.10 (-0.17, 0.37)  | D.01 0.1 10 100 Favours (Kanglaite) Favours (Chemotherapy) Risk Difference M.H, Fixed, 95% Cl           |
| Total events<br>Heterogeneity: Chi <sup>2</sup> =<br>Test for overall effect:<br>Bao H 2019<br>Chen C 2018<br>Chen V 2018<br>Chen V 2016<br>Chen V 2013<br>Guan XO 2009<br>He LT 2017<br>Huang ZB 2010<br>Jia JN 2018<br>Li HV 2017<br>Li L 2012<br>Li XD 2016<br>Liang GC 2014<br>Liu F 2019<br>Liu JQ 2011<br>Liu XH 2015<br>Liong SG 2017<br>Mu Q 2018<br>Sun SQ 2012<br>Wang CY 2018<br>Wang LY 2016<br>Wang L 2014<br>Wang L 2014  | 588<br>19.51, df<br>Z = 6.91 (<br>Kangla<br>Events<br>225<br>166<br>177<br>222<br>133<br>5<br>222<br>133<br>5<br>222<br>133<br>5<br>222<br>138<br>15<br>166<br>8<br>244<br>233<br>100<br>400<br>45<br>277<br>188<br>177<br>244<br>15<br>17<br>24<br>15<br>188<br>19<br>19<br>24<br>15<br>27<br>24<br>15<br>27<br>24<br>24<br>24<br>24<br>29<br>29<br>20<br>29<br>20<br>20<br>20<br>20<br>20<br>20<br>20<br>20<br>20<br>20<br>20<br>20<br>20   | = 31 (P < 0.0)<br>(P < 0.0)<br>31 (P < 0.0)<br>31 (P < 0.0)<br>44<br>51 (2)<br>54<br>51 (2)<br>54<br>35 (2)<br>53 (2)<br>54 (2)<br>55 | 415<br>= 0.95); /=<br>0.96); /=<br>0001)<br>Chemothin<br>Events<br>17<br>11<br>15<br>15<br>18<br>9<br>4<br>11<br>13<br>10<br>14<br>9<br>14<br>10<br>14<br>9<br>14<br>10<br>14<br>10<br>14<br>10<br>14<br>10<br>12<br>17<br>11<br>13<br>10<br>14<br>14<br>16<br>17<br>11<br>17<br>11<br>13<br>10<br>14<br>14<br>16<br>17<br>11<br>17<br>11<br>13<br>10<br>14<br>14<br>14<br>16<br>17<br>17<br>11<br>13<br>10<br>14<br>14<br>14<br>16<br>17<br>11<br>17<br>11<br>13<br>10<br>14<br>14<br>16<br>17<br>11<br>17<br>11<br>13<br>10<br>14<br>14<br>16<br>17<br>11<br>17<br>11<br>13<br>10<br>14<br>14<br>16<br>17<br>17<br>11<br>13<br>10<br>14<br>14<br>16<br>17<br>17<br>11<br>17<br>11<br>13<br>10<br>14<br>16<br>17<br>17<br>11<br>13<br>10<br>14<br>14<br>16<br>17<br>17<br>11<br>17<br>17<br>17<br>17<br>17<br>17<br>17  | = 0%.<br>Total<br>311<br>300<br>444<br>511<br>277<br>544<br>355<br>544<br>355<br>311<br>400<br>200<br>633<br>355<br>555<br>433<br>404<br>407<br>200<br>635<br>435<br>444<br>407<br>200<br>635<br>444<br>407<br>200<br>635<br>545<br>437<br>447<br>447<br>555<br>555<br>555<br>437<br>447<br>457<br>555<br>555<br>555<br>437<br>447<br>555<br>555<br>555<br>555<br>555<br>555<br>55                              | Weight<br>2.4%<br>2.3%<br>4.0%<br>2.1%<br>0.9%<br>4.2%<br>2.7%<br>3.0%<br>3.0%<br>3.0%<br>3.1%<br>4.9%<br>2.7%<br>3.3%<br>3.3%<br>3.3%<br>3.3%<br>3.3%<br>3.3%<br>3.3%<br>3                    | Risk Difference<br>M-H, Fixed, 95% CI<br>0.28 (0.03, 0.48)<br>0.17 (+0.08, 0.41)<br>0.05 (+0.16, 0.25)<br>0.08 (+0.11, 0.27)<br>0.13 (+0.13, 0.39)<br>0.08 (+0.30, 0.47)<br>0.20 (0.03, 0.37)<br>0.17 (+0.08, 0.40)<br>0.16 (+0.08, 0.40)<br>0.16 (+0.08, 0.40)<br>0.05 (+0.16, 0.26)<br>-0.01 (+0.20, 0.17)<br>0.26 (0.04, 0.47)<br>0.22 (0.05, 0.39)<br>0.09 (+0.14, 0.31)<br>0.18 (0.00, 0.36)<br>0.09 (+0.14, 0.31)<br>0.18 (0.00, 0.36)<br>0.03 (+0.18, 0.24)<br>0.21 (0.02, 0.41)<br>0.11 (+0.11, 0.37)<br>0.16 (+0.04, 0.36)<br>0.16 (+0.04, 0.36)<br>0.21 (0.12, 0.17)<br>0.16 (+0.04, 0.36)<br>0.21 (0.02, 0.41)<br>0.11 (+0.17, 0.37)<br>0.16 (+0.04, 0.36)<br>0.20 (+0.04, 0.36)<br>0.20 (+0.17, 0.37)<br>0.20 (+0.17, 0.37         | D.01 0.1 10 100 Favours (Kanglaite) Favours (Chemotherapy) Risk Difference M.H. Fixed, 95% Cl           |
| Total events<br>Heterogeneity: Chi <sup>2</sup> =<br>Test for overall effect:<br>Study or Subgroup<br>Bao H 2019<br>Chen C 2018<br>Chen V 2018<br>Chen Y 2018<br>Chen Y 2018<br>Chen Y 2018<br>Chen Y 2018<br>Chen Y 2018<br>Li HY 2017<br>Huang ZB 2010<br>Jia JN 2018<br>Li HY 2017<br>Li L 2012<br>Li XD 2016<br>Liang J 2018<br>Li XJ 2016<br>Li XJ 2016<br>Li XJ 2015<br>Long SG 2014<br>Liu XH 2015<br>Liu XH 2015<br>Long SG 2017<br>Mu Q 2018<br>Sun SQ 2012<br>Wang CY 2018<br>Wang HY 2016<br>Wang L 2014<br>Wang Y 2017  | 588 19.51, df Z= 6.91 Kangla Events 25 16 25 16 17 22 13 5 22 18 15 16 16 18 24 23 10 40 15 27 18 17 24 15 18 17 24 15 18 10 19 26 26 26 26 26 26 26 26 26 27 27 27 26 27 27 26 27 27 26 27 27 26 27 27 26 27 27 26 27 27 26 27 27 26 27 27 26 27 27 27 26 27 27 27 26 27 27 27 26 27 27 27 27 27 27 27 27 27 27 27 27 27   | = 31 (P < 0.0<br>Total 31<br>300 44<br>31 28<br>32 34<br>34 451<br>28 35<br>34 35<br>34 35<br>34 35<br>34 35<br>34 35<br>34 35<br>35 5<br>43 35<br>43 35<br>45<br>45<br>45<br>45<br>45<br>45<br>45<br>45<br>45<br>4   | 415<br>= 0.95); /*<br>00001)<br>Chemothin<br>Events<br>17<br>11<br>15<br>18<br>9<br>4<br>11<br>13<br>10<br>14<br>10<br>14<br>10<br>14<br>10<br>8<br>26<br>12<br>17<br>14<br>15<br>16<br>17<br>11<br>17<br>11<br>13<br>10<br>14<br>10<br>17<br>11<br>13<br>10<br>14<br>11<br>15<br>18<br>9<br>4<br>11<br>13<br>10<br>14<br>11<br>15<br>18<br>9<br>4<br>11<br>13<br>10<br>14<br>11<br>15<br>18<br>9<br>4<br>11<br>11<br>15<br>18<br>9<br>4<br>11<br>11<br>15<br>18<br>9<br>4<br>11<br>11<br>15<br>18<br>9<br>4<br>11<br>11<br>15<br>18<br>9<br>10<br>11<br>11<br>15<br>18<br>9<br>10<br>11<br>11<br>15<br>18<br>9<br>10<br>11<br>11<br>15<br>18<br>9<br>10<br>11<br>11<br>15<br>18<br>9<br>10<br>11<br>11<br>13<br>10<br>14<br>11<br>15<br>18<br>9<br>14<br>11<br>11<br>15<br>18<br>9<br>14<br>11<br>11<br>13<br>10<br>14<br>11<br>17<br>17<br>11<br>13<br>10<br>14<br>11<br>16<br>17<br>17<br>11<br>17<br>18<br>9<br>14<br>11<br>17<br>18<br>9<br>14<br>11<br>17<br>18<br>9<br>14<br>11<br>17<br>18<br>18<br>10<br>14<br>11<br>17<br>18<br>18<br>10<br>14<br>11<br>17<br>18<br>18<br>18<br>10<br>14<br>11<br>17<br>18<br>18<br>10<br>14<br>11<br>17<br>18<br>18<br>17<br>17<br>14<br>11<br>17<br>18<br>18<br>12<br>17<br>14<br>15<br>18<br>11<br>17<br>18<br>18<br>12<br>17<br>14<br>11<br>17<br>18<br>12<br>12<br>12<br>12<br>12<br>12<br>12<br>12<br>12<br>12 | = 0%.<br><b>Total</b><br>311<br>300<br>444<br>511<br>277<br>12<br>27<br>54<br>355<br>54<br>35<br>311<br>40<br>399<br>400<br>633<br>355<br>433<br>455<br>433<br>455<br>433<br>444<br>40<br>40<br>40<br>40<br>40<br>40<br>40<br>40<br>4   | Weight<br>2.4%<br>2.3%<br>3.4%<br>2.1%<br>0.9%<br>2.7%<br>2.4%<br>3.0%<br>3.0%<br>3.0%<br>3.0%<br>3.0%<br>3.0%<br>3.3%<br>3.2%<br>3.3%<br>3.3%<br>3.3%<br>3.3%<br>3.3%<br>3.3                  | Risk Difference<br>M-H, Fixed, 95% CI<br>0.28 (0.03, 0.48)<br>0.17 (-0.08, 0.41)<br>0.05 (-0.16, 0.25)<br>0.08 (-0.11, 0.27)<br>0.13 (-0.13, 0.39)<br>0.08 (-0.30, 0.47)<br>0.20 (0.03, 0.37)<br>0.17 (-0.06, 0.40)<br>0.16 (-0.08, 0.40)<br>0.16 (-0.16, 0.26)<br>-0.01 (-0.20, 0.17)<br>0.26 (0.04, 0.47)<br>0.26 (0.04, 0.47)<br>0.22 (0.05, 0.39)<br>0.09 (-0.14, 0.31)<br>0.18 (-0.04, 0.36)<br>0.08 (-0.11, 0.30)<br>0.3 (-0.12, 0.43)<br>0.3 (-0.12, 0.43)<br>0.3 (-0.12, 0.43)<br>0.3 (-0.14, 0.31)<br>0.3 (-0.14, 0.33)<br>0.20 (-0.14, 0.33)<br>0.21 (-0.02, 0.41)<br>0.21 (-0.02, 0.41)<br>0.21 (-0.04, 0.36)<br>0.39 (0.18, 0.60)<br>0.39 (0.18, 0.60)<br>0.30 (0.18, 0.60 | 0.01 0.1 10 100 Favours (Kanglaite) Favours (Chemotherapy) Risk Difference M.H. Fixed, 95% Cl           |
| Total events<br>Heterogeneity: Chi <sup>2</sup> =<br>Test for overall effect:<br>Study or Subgroup<br>Bao H 2019<br>Chen C 2018<br>Chen V 2018<br>Chen V 2018<br>Chen V 2018<br>Chen YZ 2003<br>Guan X0 2009<br>He LT 2017<br>Huang ZB 2010<br>Jia JN 2018<br>Li HY 2017<br>Li L 2012<br>Li XD 2016<br>Liang SG 2014<br>Li W 2015<br>Long SG 2014<br>Li W 2015<br>Long SG 2017<br>Mu Q 2018<br>Sun SQ 2012<br>Wang HY 2016<br>Wang HY 2016<br>Wang L 2014<br>Wang Y 2017<br>Yan QH 2018   | 588 19.51, df Z = 6.91 Kangla Events 25 16 17 22 13 5 25 18 15 16 8 24 23 10 10 15 27 18 10 15 18 10 18 18 10 19 26 23 23 24 23 24 24 23 24 23 24 24 24 25 25 25 25 25 25 25 25 25 25 25 25 25  | = 31 (P)<br>(P)<br>(P)<br>(P)<br>(P)<br>(P)<br>(P)<br>(P)   | 415<br>'= 0.95); /=<br>00001)<br>Chemothin<br>Events<br>17<br>11<br>15<br>18<br>9<br>4<br>11<br>13<br>10<br>14<br>13<br>10<br>14<br>13<br>10<br>14<br>10<br>14<br>10<br>14<br>15<br>14<br>15<br>14<br>15<br>14<br>11<br>17<br>26<br>12<br>17<br>14<br>15<br>14<br>11<br>17<br>26<br>12<br>17<br>14<br>11<br>17<br>13<br>18<br>19<br>19<br>19<br>19<br>19<br>19<br>19<br>19<br>19<br>19<br>19<br>19<br>19   | = 0%<br>rapy<br>Total<br>31<br>300<br>44<br>51<br>27<br>12<br>24<br>35<br>31<br>41<br>40<br>39<br>40<br>20<br>33<br>35<br>55<br>43<br>35<br>54<br>35<br>54<br>40<br>40<br>40<br>40<br>40<br>40<br>40<br>40<br>40<br>4   | Weiaht<br>2.4%<br>2.3%<br>3.4%<br>4.0%<br>2.1%<br>0.9%<br>2.4%<br>3.2%<br>3.0%<br>3.0%<br>3.0%<br>3.0%<br>3.0%<br>3.0%<br>3.0%<br>3.0  | Risk Difference<br><u>M-H, Fixed, 95% CI</u><br>0.28 (0.03, 0.48)<br>0.17 (-0.08, 0.41)<br>0.55 (-0.16, 0.25)<br>0.08 (-0.11, 0.27)<br>0.13 (-0.13, 0.39)<br>0.08 (-0.30, 0.47)<br>0.20 (0.03, 0.37)<br>0.17 (-0.06, 0.40)<br>0.15 (-0.08, 0.40)<br>0.55 (-0.16, 0.28)<br>-0.01 (-0.20, 0.17)<br>0.26 [0.04, 0.47]<br>0.25 [0.16, 0.28]<br>0.09 (-0.14, 0.31]<br>0.29 (0.02, 0.39]<br>0.09 (-0.11, 0.30]<br>0.09 (-0.11, 0.30]<br>0.09 (-0.11, 0.30]<br>0.03 (-0.18, 0.24)<br>0.11 (-0.17, 0.37]<br>0.16 (-0.04, 0.36]<br>0.39 (0.18, 0.60)<br>0.04 (-0.16, 0.24)  | D.01 0.1 10 100 Favours [Kanglaite] Favours [Chemotherapy] Risk Difference M.H. Fixed, 95% Cl           |
| Total events<br>Heterogeneity: Chi <sup>2</sup> =<br>Test for overall effect:<br>Bao H 2019<br>Chen C 2018<br>Chen V 2018<br>Chen V 2018<br>Chen V 2018<br>Chen V 2018<br>Chen YZ 2003<br>Guan XO 2009<br>He LT 2017<br>Huang ZB 2010<br>Jia JN 2018<br>Li HV 2017<br>Li L 2012<br>Li XD 2016<br>Liang J 2018<br>Liang SG 2014<br>Liu F 2015<br>Liong SG 2017<br>Mu Q 2018<br>Sun SQ 2012<br>Wang CY 2018<br>Wang LY 2016<br>Wang LY 2016<br>Wang L 2014<br>Wang L 2014<br>Wang L 2014<br>Wang L 2017<br>Yan QH 2018  | 588 19.51, df Z = 6.91 Kangla Events 25 16 17 22 13 5 22 13 5 22 18 15 16 8 24 23 10 40 15 27 18 16 8 24 23 10 40 15 27 18 18 10 10 19 24 15 18 18 10 19 24 23 13 13 13 13 14 15 18 10 19 24 23 13 13 13 14 15 18 18 10 19 24 23 23 24 23 24 24 25 25 25 25 25 25 25 25 25 25 25 25 25  | = 31 (P)<br>F doi:<br>10 - 10 - 10 - 10 - 10 - 10 - 10 - 10 -   | 415<br>(= 0.95);  =<br>00001)<br>Chemoth<br>Events<br>17<br>11<br>15<br>18<br>9<br>4<br>11<br>13<br>10<br>14<br>13<br>10<br>14<br>13<br>10<br>14<br>10<br>14<br>10<br>14<br>10<br>14<br>10<br>14<br>11<br>15<br>14<br>11<br>17<br>18<br>26<br>12<br>17<br>17<br>18<br>18<br>19<br>14<br>10<br>19<br>14<br>10<br>19<br>14<br>10<br>19<br>14<br>10<br>19<br>14<br>10<br>19<br>14<br>10<br>19<br>14<br>10<br>19<br>14<br>10<br>19<br>14<br>10<br>19<br>14<br>17<br>17<br>17<br>18<br>18<br>19<br>19<br>19<br>19<br>19<br>19<br>19<br>19<br>19<br>19<br>19<br>19<br>19   | = 0%.<br>Total<br>311<br>300<br>444<br>511<br>277<br>144<br>451<br>277<br>544<br>355<br>544<br>355<br>311<br>400<br>200<br>633<br>555<br>555<br>430<br>407<br>407<br>407<br>407<br>407<br>407<br>407<br>40  | Weight<br>2.4%<br>2.3%<br>4.0%<br>2.1%<br>2.7%<br>2.4%<br>3.2%<br>3.0%<br>3.0%<br>3.0%<br>3.1%<br>4.9%<br>4.3%<br>3.3%<br>3.3%<br>3.3%<br>3.3%<br>3.3%<br>3.3%<br>3.3                          | Risk Difference<br><u>M-H, Fixed, 95% CI</u><br>0.28 (0.03, 0.48)<br>0.17 (-0.08, 0.41)<br>0.05 (-0.16, 0.25)<br>0.08 (-0.31, 0.39)<br>0.08 (-0.30, 0.47)<br>0.20 (0.03, 0.37)<br>0.17 (-0.06, 0.40)<br>0.16 (-0.08, 0.40)<br>0.05 (-0.16, 0.26)<br>-0.01 (-0.20, 0.17)<br>0.25 (0.04, 0.47)<br>0.22 (0.05, 0.40)<br>0.03 (-0.16, 0.26)<br>0.03 (-0.26, 0.33)<br>0.09 (-0.14, 0.31)<br>0.18 (0.00, 0.36)<br>0.09 (-0.14, 0.31)<br>0.18 (0.00, 0.36)<br>0.03 (-0.18, 0.24)<br>0.21 (0.02, 0.41)<br>0.11 (-0.17, 0.37)<br>0.16 (-0.04, 0.36)<br>0.09 (-0.17, 0.37)<br>0.16 (-0.04, 0.36)<br>0.09 (-0.17, 0.37)<br>0.16 (-0.04, 0.36)<br>0.09 (-0.13, 0.30)<br>0.04 (-0.16, 0.24)<br>0.09 (-0.13, 0.30)   | D.01 0.1 10 100 Favours (Kanglaite) Favours (Chemotherapy) Risk Difference M.H. Fixed, 95% Cl           |
| Total events<br>Heterogeneity: Chi <sup>2</sup> =<br>Test for overall effect:<br>Study or Subgroup<br>Bao H 2019<br>Chen C 2018<br>Chen V 2018<br>Chen V 2018<br>Chen V 2018<br>Chen V 2018<br>Chen V 2018<br>He LT 2017<br>Huang ZB 2010<br>Jia JN 2018<br>Li HY 2017<br>Li L 2012<br>Li L 2015<br>Li L 2015<br>Li L 2015<br>Li L 2018<br>Sun SQ 2012<br>Wang CY 2018<br>Wang Y 2018<br>Wang L 2014<br>Wang Y 2017<br>Yang L 2018<br>Yang L 2018 | 588 19.51, df Z= 6.91 Kangla Events 25 16 17 22 13 5 22 18 15 16 16 16 8 24 23 10 40 15 27 18 17 24 15 18 17 24 15 18 17 24 15 18 19 26 23 13 19 26 23 13 13 19 26 23 13 13 13 13 12 14 15 18 14 15 18 18 19 26 23 13 13 13 13 13 13 13 13 13 13 13 13 14 15 15 15 15 15 15 15 15 15 15 15 15 15  | = 31 (P( 9.0.<br>Total 31<br>31 300<br>44<br>31 28<br>32 28<br>34<br>35<br>54<br>35<br>54<br>35<br>54<br>33<br>39<br>40<br>23<br>35<br>55<br>43<br>35<br>55<br>43<br>42<br>24<br>47<br>35<br>42<br>24<br>43<br>36<br>35<br>55<br>43<br>43<br>44<br>43<br>55<br>55<br>43<br>43<br>44<br>43<br>55<br>55<br>43<br>43<br>44<br>45<br>55<br>55<br>43<br>45<br>45<br>55<br>45<br>45<br>45<br>45<br>45<br>45<br>45   | 415<br>(20,95), (*<br>0001)<br>Chemothin<br>Events<br>17<br>11<br>15<br>18<br>9<br>4<br>11<br>13<br>10<br>14<br>14<br>10<br>8<br>26<br>12<br>17<br>14<br>15<br>14<br>15<br>14<br>15<br>14<br>17<br>17<br>11<br>17<br>11<br>18<br>9<br>4<br>11<br>13<br>10<br>14<br>11<br>15<br>16<br>17<br>11<br>17<br>11<br>18<br>9<br>4<br>11<br>17<br>11<br>18<br>9<br>4<br>11<br>17<br>11<br>18<br>9<br>4<br>11<br>17<br>11<br>18<br>9<br>4<br>11<br>17<br>11<br>18<br>9<br>4<br>11<br>17<br>11<br>18<br>9<br>4<br>11<br>17<br>18<br>19<br>11<br>11<br>15<br>18<br>9<br>4<br>11<br>11<br>15<br>18<br>9<br>10<br>11<br>11<br>15<br>18<br>9<br>10<br>11<br>11<br>15<br>18<br>9<br>10<br>11<br>11<br>13<br>10<br>14<br>11<br>15<br>16<br>17<br>11<br>11<br>15<br>18<br>9<br>11<br>11<br>13<br>10<br>8<br>26<br>17<br>17<br>17<br>17<br>17<br>11<br>18<br>9<br>17<br>11<br>18<br>9<br>10<br>11<br>14<br>15<br>17<br>17<br>17<br>17<br>17<br>17<br>17<br>18<br>9<br>17<br>17<br>17<br>18<br>9<br>10<br>14<br>11<br>17<br>17<br>17<br>17<br>17<br>17<br>17<br>17<br>17   | = 0%.<br>Total<br>311<br>300<br>444<br>511<br>277<br>122<br>544<br>355<br>544<br>349<br>400<br>633<br>3555<br>433<br>455<br>433<br>464<br>400<br>633<br>355<br>433<br>444<br>407<br>407<br>407<br>407<br>407<br>407<br>407  | Weight<br>2.4%<br>2.3%<br>3.4%<br>4.0%<br>2.1%<br>0.9%<br>2.4%<br>3.0%<br>3.0%<br>3.0%<br>3.1%<br>3.1%<br>3.3%<br>3.3%<br>3.3%<br>3.3%<br>3.3%<br>3.3  | Risk Difference<br>M-H, Fixed, 95% CI<br>0.28 (0.03, 0.48)<br>0.17 (-0.08, 0.41)<br>0.05 (-0.16, 0.25)<br>0.08 (-0.11, 0.27)<br>0.13 (-0.13, 0.39)<br>0.08 (-0.30, 0.47)<br>0.20 (0.03, 0.37)<br>0.17 (-0.06, 0.40)<br>0.16 (-0.08, 0.40)<br>0.05 (-0.16, 0.26)<br>-0.01 (-0.20, 0.17)<br>0.26 (0.04, 0.47)<br>0.22 (0.05, 0.39)<br>0.03 (-0.26, 0.33)<br>0.22 (0.05, 0.39)<br>0.03 (-0.26, 0.33)<br>0.22 (0.05, 0.39)<br>0.09 (-0.14, 0.31)<br>0.09 (-0.14, 0.31)<br>0.09 (-0.14, 0.31)<br>0.03 (-0.18, 0.24)<br>0.21 (-0.2, 0.41)<br>0.11 (-0.11, 0.34)<br>0.02 (-0.19, 0.23)<br>0.04 (-0.16, 0.24)<br>0.09 (-0.18, 0.60)<br>0.04 (-0.16, 0.34)<br>0.09 (-0.18, 0.60)<br>0.04 (-0.16, 0.34)<br>0.09 (-0.13, 0.30)<br>0.09 (-0.16, 0.34)<br>0.09 (-0.13, 0.30)<br>0.09 (-0.16, 0.34)<br>0.09 (-0.16, 0.34)<br>0.09 (-0.16, 0.34)<br>0.09 (-0.13, 0.30)<br>0.09 (-0.16, 0.34)<br>0.09 (-0.16, 0.34)<br>0.09 (-0.16, 0.34)<br>0.09 (-0.16, 0.34)<br>0.09 (-0.13, 0.30)<br>0.09 (-0.16, 0.34)<br>0.09 (-0.16, 0.34)<br>0.09 (-0.16, 0.34)<br>0.09 (-0.16, 0.34)<br>0.09 (-0.16, 0.34)<br>0.09 (-0.16, 0.34)<br>0.09 (-0.13, 0.30)<br>0.09 (-0.16, 0.34)<br>0.09 (-0.16, 0.34)<br>0.09 (-0.16, 0.34)<br>0.09 (-0.16, 0.34)<br>0.09 (-0.13, 0.30)<br>0.09 (-0.16, 0.34)<br>0.09 (-0.16, 0.         | D.01 0.1 10 100 Favours (Kanglaite) Favours (Chemotherapy) Risk Difference M.H. Fixed, 95% Cl           |
| Total events<br>Heterogeneity: Chi <sup>2</sup> =<br>Test for overall effect:<br>Study or Subgroup<br>Bao H 2019<br>Chen C 2018<br>Chen V 2018<br>Chen V 2018<br>Chen V 2018<br>Chen Y 2003<br>Guan X0 2009<br>He LT 2017<br>Huang ZB 2010<br>Jia JN 2018<br>Li HY 2017<br>Li L 2012<br>Li L 2012<br>Li L 2012<br>Li L 2012<br>Li L 2016<br>Liang SG 2014<br>Liu F 2019<br>Liu JQ 2016<br>Liu XH 2015<br>Long SG 2017<br>Mu Q 2018<br>Sun SQ 2012<br>Wang LY 2016<br>Wang L 2014<br>Wang Y 2017<br>Yan GH 2018<br>Yang J 2018<br>Yang J 2017<br>Yan GH 2018<br>Yang J 2016<br>Yang SJ 2003<br>Yao J 2017  | 588 19.51, df Z= 6.91 Kangla Events 25 16 25 16 17 22 13 5 22 18 15 16 16 8 24 23 10 10 15 27 18 17 24 15 18 10 19 26 23 10 10 19 26 23 10 19 19 26 23 10 19 19 26 23 10 19 19 26 23 10 19 19 26 23 10 19 19 26 23 10 19 19 26 23 10 19 19 26 23 10 19 19 26 23 10 19 19 26 23 10 19 26 23 10 19 26 23 10 19 26 23 10 19 26 23 10 19 26 23 10 19 26 23 10 19 26 23 10 19 26 23 10 19 26 23 10 19 26 23 10 19 26 23 10 19 26 23 10 19 26 23 10 19 26 23 10 19 26 23 10 19 26 23 10 19 26 23 10 19 26 23 10 19 26 23 19 26 23 10 19 26 23 19 19 26 23 19 19 26 23 19 19 26 23 19 19 26 23 19 19 26 23 10 19 26 23 19 19 26 23 19 26 23 19 26 23 23 2 23 2 2 2 2 2 2 2 2 2 2 2 2 2   | = 31 (P < 0.0<br>P < 0.0<br>Total 31<br>300 44<br>51 28<br>35 31<br>32 35<br>331 42<br>35 331<br>39 40<br>35 55<br>33 42<br>35 55<br>35 55<br>35 55<br>35 55<br>35 55<br>32 42<br>47<br>47<br>47<br>47<br>47<br>47<br>47<br>47<br>47<br>47<br>47<br>47<br>47  | 415<br>'= 0.95); /=<br>00001)<br>Chemothin<br>Events<br>17<br>11<br>15<br>5<br>18<br>9<br>4<br>11<br>13<br>10<br>14<br>13<br>10<br>14<br>10<br>8<br>26<br>12<br>17<br>14<br>15<br>14<br>10<br>8<br>26<br>12<br>17<br>17<br>11<br>13<br>10<br>10<br>10<br>10<br>10<br>10<br>10<br>10<br>10<br>10<br>10<br>10<br>10  | = 0%<br>trapy<br>Total<br>31<br>300<br>44<br>451<br>27<br>712<br>27<br>54<br>35<br>54<br>35<br>311<br>40<br>39<br>9<br>40<br>63<br>35<br>55<br>43<br>35<br>43<br>44<br>40<br>40<br>43<br>55<br>44<br>40<br>40<br>40<br>40<br>40<br>40<br>40<br>40<br>40   | Weight<br>2.4%<br>2.3%<br>3.4%<br>4.0%<br>2.1%<br>0.9%<br>2.4%<br>3.0%<br>3.0%<br>3.0%<br>3.0%<br>3.0%<br>3.0%<br>3.0%<br>3.0  | Risk Difference<br>M-H, Fixed, 95% CI<br>0.28 (0.03, 0.48)<br>0.17 (-0.08, 0.41)<br>0.55 (-0.16, 0.25)<br>0.08 (-0.11, 0.27)<br>0.13 (-0.13, 0.39)<br>0.08 (-0.30, 0.47)<br>0.20 (0.03, 0.37)<br>0.17 (-0.06, 0.40)<br>0.15 (-0.16, 0.28)<br>-0.01 (-0.20, 0.17)<br>0.26 (0.04, 0.47)<br>0.26 (0.04, 0.47)<br>0.26 (0.04, 0.47)<br>0.26 (0.04, 0.47)<br>0.26 (0.04, 0.47)<br>0.22 (0.05, 0.38)<br>0.09 (-0.14, 0.31)<br>0.18 (0.00, 0.36)<br>0.08 (-0.11, 0.30)<br>0.08 (-0.11, 0.30)<br>0.03 (-0.18, 0.24)<br>0.21 (0.02, 0.41)<br>0.21 (0.02, 0.41)<br>0.29 (0.18, 0.60)<br>0.39 (0.18, 0.60)<br>0.39 (0.18, 0.60)<br>0.40 (-0.16, 0.34)<br>0.99 (-0.16, 0.34)<br>0.       | D.01 0.1 10 100 Favours [Kanglaite] Favours [Chemotherapy] Risk Difference M.H. Fixed, 95% Cl           |
| Total events<br>Heterogeneity: Chi <sup>2</sup> =<br>Test for overall effect:<br>Bao H 2019<br>Chen C 2018<br>Chen V 2018<br>Chen V 2018<br>Chen V 2018<br>Chen V 2018<br>Chen YZ 2003<br>Guan X0 2009<br>He LT 2017<br>Huang ZB 2010<br>Jia JN 2018<br>Li HY 2017<br>Li L 2012<br>Li XD 2016<br>Liang SG 2014<br>Liu F 2015<br>Long SG 2017<br>Mu Q 2018<br>Sun SQ 2012<br>Wang CY 2018<br>Wang HY 2016<br>Wang L 2014<br>Wang Y 2017<br>Yan QH 2018<br>Yang L 2018<br>Yang L 2018<br>Yang L 2018<br>Yang L 2018<br>Yang L 2018<br>Yang J 2013<br>Yang J 2013<br>Yang J 2013   | 588 19.51, df Z= 6.91 Kangla Events 25 16 17 22 13 5 16 17 22 18 15 16 8 24 23 10 10 15 27 18 10 15 27 18 10 15 27 18 10 19 24 15 18 10 19 24 23 13 9 19 22 22 22 22 22 22 22 22 22 22 22 22 22   | = 31 (P)<br>= 31 (P)<br>Total 2<br>31 30<br>30<br>44<br>51<br>28<br>31<br>28<br>32<br>31<br>28<br>35<br>31<br>38<br>39<br>40<br>23<br>35<br>55<br>55<br>55<br>55<br>55<br>55<br>55<br>55<br>5   | 415<br>= 0.95); /*<br>00001)<br>Chemothic<br>Events<br>17<br>11<br>15<br>18<br>9<br>4<br>17<br>11<br>15<br>18<br>9<br>4<br>11<br>13<br>10<br>14<br>10<br>14<br>10<br>14<br>15<br>14<br>15<br>14<br>15<br>12<br>12<br>12<br>12<br>12<br>12<br>12<br>12<br>12<br>12  | = 0%<br>trapy<br>Total<br>31<br>300<br>44<br>51<br>27<br>12<br>24<br>35<br>31<br>41<br>40<br>39<br>40<br>20<br>33<br>35<br>55<br>54<br>33<br>40<br>40<br>20<br>20<br>40<br>40<br>40<br>40<br>40<br>40<br>40<br>40<br>40<br>4  | Weiaht<br>2.4%<br>2.3%<br>3.4%<br>4.0%<br>2.1%<br>0.9%<br>2.4%<br>3.2%<br>3.0%<br>3.0%<br>3.0%<br>3.0%<br>3.0%<br>3.0%<br>3.0%<br>3.0  | Risk Difference<br><u>M-H, Fixed, 95% CI</u><br>0.28 (0.03, 0.48)<br>0.17 (-0.08, 0.41)<br>0.55 (-0.13, 0.39)<br>0.08 (-0.11, 0.27)<br>0.13 (-0.13, 0.39)<br>0.08 (-0.30, 0.47)<br>0.20 (0.03, 0.37)<br>0.17 (-0.06, 0.40)<br>0.16 (-0.08, 0.40)<br>0.55 (-0.16, 0.28)<br>-0.01 (-0.20, 0.17)<br>0.26 (0.04, 0.47)<br>0.25 (0.16, 0.28)<br>-0.01 (-0.20, 0.17)<br>0.26 (0.04, 0.47)<br>0.25 (0.16, 0.28)<br>-0.01 (-0.20, 0.17)<br>0.26 (0.04, 0.47)<br>0.25 (0.16, 0.28)<br>0.09 (-0.11, 0.30)<br>0.09 (-0.11, 0.30)<br>0.09 (-0.11, 0.34)<br>0.02 (-0.19, 0.23)<br>0.16 (-0.04, 0.36)<br>0.39 (0.18, 0.60)<br>0.04 (-0.16, 0.34)<br>0.09 (-0.13, 0.30)<br>0.09 (-0.16, 0.34)<br>0.09 (-0.16, 0.34)<br>0.05 (-0.10, 0.19)<br>0.03 (-0.19, 0.24)<br>0.15 (-0.14, 0.31)<br>0.35 (-0.19, 0.24)<br>0.35 (-0.19, 0.24)<br>0.35 (-0.10, 0.19)<br>0.33 (-0.19, 0.24)<br>0.35 (-0.10, 0.19)<br>0.33 (-0.19, 0.24)<br>0.55 (-0.10, 0.19)<br>0.35 (-0.10, 0.19)<br>0.35 (-0.10, 0.19)<br>0.35 (-0.10, 0.19)<br>0.35 (-0.10, 0.24)<br>0.55 (-0.10, 0.19)<br>0.55 (-0.10, 0.24)<br>0.55 (-0.10, 0.19)<br>0.55 (-0.10,     | D.01 0.1 10 100 Favours (Kanglaite) Favours (Chemotherapy) Risk Difference M-H, Fixed, 95% Cl           |
| Total events<br>Heterogeneity: Chi <sup>2</sup> =<br>Test for overall effect:<br>Bao H 2019<br>Chen C 2018<br>Chen V 2018<br>Chen V 2018<br>Chen V 2018<br>Chen V 2018<br>Chen YZ 2003<br>Guan XO 2009<br>He LT 2017<br>Huang ZB 2010<br>Jia JN 2018<br>Li HY 2017<br>Li L 2012<br>Li XD 2018<br>Liang J 2018<br>Liang SG 2014<br>Liu F 2019<br>Liu JQ 2011<br>Liu Y 2015<br>Liong SG 2017<br>Mu Q 2018<br>Sun SQ 2012<br>Wang CY 2018<br>Wang L 2014<br>Wang CY 2018<br>Wang L 2014<br>Wang L 2014<br>Wang L 2014<br>Wang L 2018<br>Yang L 2018<br>Yang L 2018<br>Yang J 2017<br>Ya CH 2018<br>Yang SJ 2003<br>Yao J 2017  | 588 19.51, df Z = 6.91 Kangla Events 25 16 17 22 13 5 25 16 17 22 18 15 16 8 24 23 10 40 15 27 18 18 10 10 19 24 15 18 18 10 10 19 24 15 18 18 10 10 19 24 23 13 9 19 24 23 13 13 9 19 24 23 13 13 9 19 24 23 15 18 18 10 10 19 24 23 13 13 13 13 13 14 15 18 18 10 19 24 23 13 13 14 15 18 18 18 18 18 18 18 18 18 18 18 18 18   | = 31 (P)<br>F define the second   | 415<br>(= 0.95); /=<br>(0001)<br>Chemothin<br>Events<br>17<br>11<br>15<br>18<br>9<br>4<br>11<br>13<br>10<br>14<br>9<br>14<br>9<br>14<br>10<br>14<br>10<br>14<br>10<br>14<br>10<br>12<br>17<br>11<br>13<br>10<br>14<br>10<br>14<br>10<br>12<br>17<br>11<br>13<br>10<br>14<br>10<br>11<br>13<br>10<br>14<br>10<br>11<br>13<br>10<br>14<br>10<br>11<br>13<br>10<br>11<br>13<br>10<br>14<br>10<br>11<br>13<br>10<br>11<br>13<br>10<br>14<br>10<br>11<br>13<br>10<br>14<br>10<br>11<br>13<br>10<br>14<br>10<br>11<br>13<br>10<br>14<br>10<br>11<br>13<br>10<br>11<br>13<br>10<br>14<br>10<br>11<br>13<br>10<br>11<br>13<br>10<br>14<br>10<br>11<br>11<br>12<br>12<br>12<br>12<br>12<br>12<br>12<br>12   | = 0%.<br>Total<br>311<br>300<br>344<br>511<br>277<br>544<br>355<br>544<br>355<br>311<br>400<br>200<br>633<br>440<br>200<br>635<br>555<br>433<br>440<br>200<br>635<br>555<br>433<br>440<br>400<br>200<br>635<br>555<br>436<br>437<br>400<br>400<br>400<br>400<br>400<br>400<br>400<br>40   | Weight<br>2.4%<br>2.3%<br>4.0%<br>2.1%<br>2.7%<br>2.4%<br>3.2%<br>3.0%<br>3.0%<br>3.1%<br>4.2%<br>4.3%<br>3.3%<br>3.3%<br>3.3%<br>3.3%<br>3.3%<br>3.3%<br>3.3                                  | Risk Difference<br><u>M-H, Fixed, 95% CI</u><br>0.28 [0.03, 0.48]<br>0.17 [-0.08, 0.41]<br>0.05 [-0.16, 0.25]<br>0.08 [-0.11, 0.27]<br>0.13 [-0.13, 0.39]<br>0.08 [-0.30, 0.47]<br>0.20 [0.03, 0.37]<br>0.17 [-0.06, 0.40]<br>0.05 [-0.16, 0.26]<br>-0.01 [-0.20, 0.17]<br>0.25 [0.04, 0.47]<br>0.26 [0.04, 0.40]<br>0.05 [-0.16, 0.26]<br>-0.01 [-0.20, 0.17]<br>0.26 [0.04, 0.47]<br>0.22 [0.05, 0.39]<br>0.09 [-0.14, 0.31]<br>0.18 [-0.00, 0.36]<br>0.09 [-0.14, 0.31]<br>0.18 [-0.00, 0.36]<br>0.09 [-0.14, 0.31]<br>0.18 [-0.00, 0.36]<br>0.09 [-0.14, 0.31]<br>0.18 [-0.02, 0.41]<br>0.11 [-0.17, 0.37]<br>0.16 [-0.04, 0.36]<br>0.39 [-0.18, 0.24]<br>0.39 [-0.13, 0.30]<br>0.09 [-0.13, 0.30]<br>0.09 [-0.13, 0.30]<br>0.09 [-0.14, 0.31]<br>0.03 [-0.19, 0.24]<br>0.05 [-0.10, 0.19]<br>0.03 [-0.19, 0.24]<br>0.15 [-0.01, 0.19]<br>0.03 [-0.19, 0.24]<br>0.15 [-0.01, 0.19]<br>0.19 [-0.13, 0.30]<br>0.19 [-0.14, 0.24]<br>0.15 [-0.01, 0.19]<br>0.33 [-0.19, 0.24]<br>0.15 [-0.01, 0.21]<br>0.19 [-0.14, 0.24]<br>0.15 [-0.01, 0.24]<br>0.19 [-0.14, 0.24]<br>0.15 [-0.01, 0.24]<br>0.19 [-0.14, 0.24]<br>0.25 [-0.10, 0.19]<br>0.24 [-0.14, 0.24]<br>0.25 [-0.10, 0.24]<br>0.25 [-0.10    | D.01 0.1 10 100 Favours (Kanglaite) Favours (Chemotherapy) Risk Difference M-H, Fixed, 95% Cl           |
| Total events<br>Heterogeneity: Chi <sup>2</sup> =<br>Test for overall effect:<br>Study or Subgroup<br>Bao H 2019<br>Chen C 2018<br>Chen V 2018<br>Chen V 2018<br>Chen V 2018<br>Chen Y 2003<br>Guan X0 2009<br>He LT 2017<br>Huang ZB 2010<br>Jia JN 2018<br>Li HY 2017<br>Li L 2012<br>Li XD 2016<br>Liang SG 2014<br>Li W 2015<br>Long SG 2014<br>Liu Y 2015<br>Long SG 2017<br>Mu Q 2018<br>Sun SQ 2012<br>Wang HY 2016<br>Wang HY 2016<br>Wang L 2018<br>Wang HY 2016<br>Wang L 2018<br>Wang HY 2016<br>Wang L 2014<br>Wang Y 2017<br>Yan QH 2018<br>Yang J 2013<br>Yao J 2017<br>Ye CY 2019<br>Yu T 2015<br>Zhang MM 2019  | 588 19.51, df Z= 6.91 Kangla Events 25 16 17 22 13 5 22 18 15 16 8 24 23 10 10 15 27 18 10 15 27 18 10 10 19 26 23 13 9 19 22 23 15   | = 31 (P. P. S.  | 415<br>= 0.95); /*<br>00001)<br>Chemothic<br>Events<br>17<br>11<br>15<br>18<br>9<br>4<br>17<br>11<br>15<br>18<br>9<br>4<br>11<br>13<br>10<br>14<br>10<br>14<br>15<br>14<br>15<br>12<br>12<br>12<br>12<br>12<br>12<br>12<br>12<br>15<br>15<br>21<br>16<br>16<br>16<br>17<br>17<br>11<br>15<br>18<br>9<br>4<br>17<br>11<br>13<br>10<br>14<br>10<br>14<br>10<br>17<br>11<br>15<br>18<br>9<br>4<br>17<br>11<br>13<br>10<br>14<br>10<br>14<br>10<br>14<br>10<br>17<br>11<br>15<br>18<br>9<br>4<br>11<br>13<br>10<br>14<br>10<br>14<br>10<br>14<br>10<br>14<br>10<br>14<br>10<br>14<br>10<br>14<br>10<br>14<br>10<br>14<br>10<br>10<br>14<br>10<br>10<br>10<br>10<br>10<br>10<br>10<br>10<br>10<br>11<br>13<br>10<br>14<br>10<br>14<br>10<br>12<br>17<br>11<br>13<br>10<br>14<br>10<br>14<br>10<br>14<br>10<br>14<br>15<br>14<br>15<br>14<br>15<br>16<br>16<br>16<br>16<br>16<br>16<br>16<br>16<br>16<br>16  | = 0%<br>trapy<br>Total<br>31<br>300<br>44<br>51<br>27<br>12<br>24<br>35<br>53<br>31<br>41<br>40<br>39<br>40<br>20<br>63<br>35<br>55<br>55<br>43<br>40<br>40<br>25<br>43<br>40<br>40<br>40<br>40<br>40<br>40<br>40<br>40<br>40<br>40   | Weight<br>2.4%<br>2.3%<br>3.4%<br>4.0%<br>2.1%<br>0.9%<br>2.4%<br>3.2%<br>3.0%<br>3.0%<br>3.0%<br>3.0%<br>3.0%<br>3.0%<br>3.0%<br>3.3%<br>3.3  | Risk Difference<br><u>M-H, Fixed, 95% CI</u><br>0.28 (0.03, 0.48)<br>0.17 (-0.08, 0.41)<br>0.55 (-0.16, 0.25)<br>0.08 (-0.11, 0.27)<br>0.13 (-0.13, 0.39)<br>0.08 (-0.30, 0.47)<br>0.20 (0.03, 0.37)<br>0.17 (-0.06, 0.40)<br>0.15 (-0.08, 0.40)<br>0.55 (-0.16, 0.28)<br>-0.01 (-0.20, 0.17)<br>0.26 (0.04, 0.47)<br>0.26 (0.04, 0.47)<br>0.26 (0.04, 0.47)<br>0.22 (0.05, 0.39)<br>0.09 (-0.14, 0.31)<br>0.03 (-0.18, 0.24)<br>0.03 (-0.18, 0.24)<br>0.02 (-0.19, 0.23)<br>0.10 (-0.17, 0.37)<br>0.16 (-0.04, 0.36)<br>0.39 (0.18, 0.680)<br>0.04 (-0.16, 0.24)<br>0.09 (-0.13, 0.30)<br>0.09 (-0.13, 0.30)<br>0.09 (-0.13, 0.30)<br>0.09 (-0.13, 0.30)<br>0.09 (-0.13, 0.30)<br>0.09 (-0.13, 0.30)<br>0.03 (-0.19, 0.24)<br>0.05 (-0.10, 0.19)<br>0.03 (-0.19, 0.24)<br>0.15 (-0.01, 0.31)<br>0.14 (0.02, 0.34)<br>0.15 (-0.01, 0.31)<br>0.14 (0.02, 0.34)<br>0.15 (-0.01, 0.31)<br>0.15 (-0.01, 0    | D.01 0.1 10 100 Favours (Kanglaite) Favours (Chemotherapy) Risk Difference M-H, Fixed, 95% Cl           |
| Total events<br>Heterogeneity: Chi <sup>2</sup> =<br>Test for overall effect:<br>Study or Subgroup<br>Bao H 2019<br>Chen C 2018<br>Chen V 2018<br>Chen V 2018<br>Chen V 2018<br>Chen YZ 2003<br>Guan X0 2009<br>He LT 2017<br>Huang ZB 2010<br>Jia JN 2018<br>Li HY 2017<br>Li L 2012<br>Li XD 2018<br>Liang SG 2014<br>Liu F 2019<br>Liu JQ 2011<br>Liu XD 2016<br>Liang SG 2014<br>Liu F 2015<br>Long SG 2017<br>Mu Q 2018<br>Sun SG 2012<br>Wang CY 2018<br>Wang L 2014<br>Wang Y 2017<br>Yan QH 2018<br>Yang L 2016<br>Yang SJ 2003<br>Yao J 2017<br>Ye CY 2019<br>Yu T 2015<br>Zhang MM 2019<br>Total (95% CI)   | 588 19.51, df Z = 6.91 Kangla Events 25 16 17 22 13 5 25 16 17 22 13 5 26 15 16 8 24 23 10 10 15 27 18 15 16 8 24 23 10 10 19 26 23 13 9 19 26 23 15 5 5 5 5 5 5 5 5 5 5 5 5 5 5 5 5 5 5  | = 31 (P)<br>F of Content<br>Total<br>31<br>30<br>31<br>30<br>44<br>12<br>54<br>54<br>31<br>32<br>54<br>33<br>39<br>40<br>23<br>39<br>40<br>23<br>35<br>55<br>55<br>55<br>55<br>55<br>55<br>55<br>55<br>5  | 415<br>= 0.95); F<br>00001)<br>Chemothic<br>Events<br>17<br>11<br>15<br>18<br>9<br>4<br>17<br>11<br>15<br>18<br>9<br>4<br>11<br>13<br>10<br>14<br>10<br>14<br>10<br>14<br>10<br>14<br>10<br>12<br>17<br>11<br>15<br>18<br>9<br>4<br>17<br>11<br>15<br>18<br>9<br>4<br>17<br>11<br>15<br>18<br>9<br>4<br>17<br>11<br>15<br>18<br>9<br>4<br>17<br>11<br>15<br>18<br>19<br>4<br>17<br>11<br>15<br>18<br>10<br>17<br>11<br>15<br>18<br>10<br>17<br>11<br>15<br>18<br>10<br>10<br>14<br>10<br>10<br>10<br>10<br>10<br>10<br>10<br>10<br>10<br>10  | = 0%.<br>Total<br>311<br>300<br>444<br>511<br>277<br>544<br>355<br>311<br>400<br>200<br>633<br>555<br>555<br>420<br>430<br>400<br>400<br>400<br>200<br>633<br>555<br>555<br>430<br>440<br>400<br>200<br>635<br>555<br>430<br>440<br>400<br>200<br>635<br>555<br>555<br>430<br>400<br>400<br>400<br>400<br>400<br>400<br>400   | Weight<br>2.4%<br>2.3%<br>4.0%<br>2.1%<br>2.4%<br>2.7%<br>2.4%<br>3.0%<br>3.0%<br>3.0%<br>3.1%<br>4.8%<br>3.3%<br>3.3%<br>3.3%<br>3.3%<br>3.3%<br>3.3%<br>3.3%<br>3                            | Risk Difference<br><u>M-H, Fixed, 95% CI</u><br>0.28 [0.03, 0.48]<br>0.17 [-0.08, 0.41]<br>0.05 [-0.16, 0.25]<br>0.08 [-0.1, 0.27]<br>0.13 [-0.13, 0.39]<br>0.08 [-0.30, 0.47]<br>0.20 [0.03, 0.37]<br>0.17 [-0.06, 0.40]<br>0.05 [-0.16, 0.26]<br>-0.01 [-0.20, 0.17]<br>0.25 [0.04, 0.47]<br>0.25 [0.04, 0.47]<br>0.25 [0.04, 0.47]<br>0.25 [0.04, 0.47]<br>0.25 [0.04, 0.47]<br>0.32 [0.12, 0.53]<br>0.09 [-0.14, 0.31]<br>0.18 [0.00, 0.36]<br>0.09 [-0.14, 0.31]<br>0.18 [0.00, 0.36]<br>0.09 [-0.14, 0.31]<br>0.18 [0.00, 0.36]<br>0.09 [-0.14, 0.31]<br>0.16 [-0.44, 0.35]<br>0.09 [-0.14, 0.31]<br>0.16 [-0.44, 0.35]<br>0.39 [0.18, 0.24]<br>0.21 [-0.23, 0.30]<br>0.09 [-0.13, 0.30]<br>0.09 [-0.13, 0.30]<br>0.09 [-0.16, 0.34]<br>0.03 [-0.19, 0.24]<br>0.15 [-0.01, 0.31]<br>0.18 [0.02, 0.34]<br>0.13 [0.10, 0.17]   | 0.01 0.1 10 100 Favours (Kanglaite) Favours (Chemotherapy) Risk Difference M-H, Fixed, 95% Cl           |

Figure 3 The analysis of ORR between two groups. ORR, objective response rate.

|  | Kangla   | aite   | Chemoth   | erapy   |   | Risk Ratio   |      | Risk Ratio   |    |
|--|--|--|---|---|---|--|------|--|----|
| Study or Subgroup  | Events   | Total  | Events  | Total   | Weight  | M-H, Random, 95% Cl  | 1    | M-H, Random, 95% Cl  |    |
| Bao H 2019   | 2  | 31   | 7   | 31  | 4.1%  | 0.29 (0.06, 1.27)  |      |  |    |
| Guan XQ 2009   | 6  | 12   | 10  | 12  | 12.8%   | 0.60 (0.32, 1.12)  |      |  |    |
| Jia JN 2018  | 13   | 31   | 27  | 31  | 16.6%   | 0.49 [0.31, 0.74]  |      |  |    |
| Liu JQ 2011  | 17   | 35   | 24  | 35  | 17.2%   | 0.71 (0.47, 1.07)  |      |  |    |
| Wang L 2014  | 5  | 35   | 10  | 35  | 7.8%  | 0.50 [0.19, 1.31]  |      | Contraction of the second seco |    |
| Wang Y 2017  | 1  | 35   | 3   | 32  | 2.0%  | 0.30 (0.03, 2.78)  |      |  |    |
| Yao J 2017   | 55   | 70   | 60  | 67  | 22.4%   | 0.88 (0.75, 1.02)  |      |  |    |
| Ye CY 2019   | 9  | 40   | 22  | 40  | 12.5%   | 0.41 [0.22, 0.78]  |      |  |    |
| Zhang MM 2019  | 3  | 50   | 5   | 50  | 4.6%  | 0.60 (0.15, 2.38)  |      |  |    |
| Total (95% CI)   |  | 339  |   | 333   | 100.0%  | 0.58 [0.42, 0.81]  |      | •  |    |
| Total events   | 111  |  | 168   |   | 0.26  | Conservation 6   |      |  |    |
| Heterogeneity: Tau <sup>2</sup> =  | = 0.12; Ch   | = 22   | 24. df = 3 (  | P=0.00  | 4); $l^2 = 64$  | %  | -    | - t - t  | -  |
| Test for overall effect  | Z= 3.20  | (P = 0.0   | 001)  |   |   |  | 0.01 | 6.1 1 10 11<br>Favours [Kanglalle] Favours [Chemolherapy]  | 00 |
|  | Kangla   | aite   | Chemoth   | erapy   |   | Risk Difference  |      | Risk Difference  |    |
| Study or Subgroup  | Events   | Total  | Events  | Total   | Weight  | M-H, Random, 95% Cl  | 1    | M-H, Random, 95% Cl  |    |
| Bao H 2019   | 2  | 31   | 7   | 31  | 11 5%   | -0 16 -0 33 0.011  |      |  |    |
| 0.111100.0000  |  |  |   |   | 11.00.00  | 0.1010.0010.0101   |      |  |    |
| Guan XQ 2009   | 6  | 12   | 10  | 12  | 4.9%  | -0.33 [-0.69, 0.02]  |      |  |    |
| Jia JN 2018  | 6<br>13  | 12<br>31   | 10<br>27  | 12<br>31  | 4.9%<br>9.5%  | -0.33 [-0.69, 0.02]<br>-0.45 [-0.66, -0.24]  |      |  |    |
| Jia JN 2018<br>Liu JO 2011   | 6<br>13<br>17  | 12<br>31<br>35   | 10<br>27<br>24  | 12<br>31<br>35  | 4.9%<br>9.5%<br>8.8%  | -0.33 [-0.69, 0.02]<br>-0.45 [-0.66, -0.24]<br>-0.20 [-0.43, 0.03]   |      |  |    |
| Jia JN 2018<br>Liu JQ 2011<br>Wang L 2014  | 6<br>13<br>17<br>5   | 12<br>31<br>35<br>35   | 10<br>27<br>24<br>10  | 12<br>31<br>35<br>35  | 4.9%<br>9.5%<br>8.8%<br>10.5%   | -0.33 [-0.69, 0.02]<br>-0.45 [-0.66, -0.24]<br>-0.20 [-0.43, 0.03]<br>-0.14 [-0.33, 0.05]  |      |  |    |
| Jia JN 2018<br>Liu JQ 2018<br>Wang L 2014<br>Wang Y 2017   | 6<br>13<br>17<br>5   | 12<br>31<br>35<br>35<br>35                                   | 10<br>27<br>24<br>10<br>3   | 12<br>31<br>35<br>35<br>32                                    | 4.9%<br>9.5%<br>8.8%<br>10.5%<br>14.9%  | -0.33 [-0.69, 0.02]<br>-0.45 [-0.66, -0.24]<br>-0.20 [-0.43, 0.03]<br>-0.14 [-0.33, 0.05]<br>-0.07 [-0.18, 0.05]   |      |  |    |
| Jia JN 2018<br>Liu JQ 2011<br>Wang L 2014<br>Wang Y 2017<br>Yao J 2017   | 6<br>13<br>17<br>5<br>1<br>55                                | 12<br>31<br>35<br>35<br>35<br>35<br>70                       | 10<br>27<br>24<br>10<br>3<br>60                                   | 12<br>31<br>35<br>35<br>32<br>67                              | 4.9%<br>9.5%<br>8.8%<br>10.5%<br>14.9%<br>14.5%   | -0.33 [-0.69, 0.02]<br>-0.45 [-0.66, -0.24]<br>-0.20 [-0.43, 0.03]<br>-0.14 [-0.33, 0.05]<br>-0.07 [-0.18, 0.05]<br>-0.11 [-0.23, 0.01]  |      |  |    |
| Guan XG 2009<br>Jia JN 2018<br>Liu JG 2011<br>Wang L 2014<br>Wang Y 2017<br>Yao J 2017<br>Ye CY 2019   | 6<br>13<br>17<br>5<br>1<br>55<br>9                           | 12<br>31<br>35<br>35<br>35<br>70<br>40                       | 10<br>27<br>24<br>10<br>3<br>60<br>22                             | 12<br>31<br>35<br>35<br>32<br>67<br>40                        | 4.9%<br>9.5%<br>8.8%<br>10.5%<br>14.9%<br>14.5%<br>9.9%   | -0.33 [-0.69, 0.02]<br>-0.45 [-0.66, -0.24]<br>-0.20 [-0.43, 0.03]<br>-0.14 [-0.33, 0.05]<br>-0.07 [-0.18, 0.05]<br>-0.11 [-0.23, 0.01]<br>-0.33 [-0.53, -0.12]  |      |  |    |
| Guan Xu 2009<br>Jia JN 2018<br>Liu Jû 2011<br>Wang L 2014<br>Wang Y 2017<br>Yao J 2017<br>Ye CY 2019<br>Zhang MM 2019  | 6<br>13<br>17<br>5<br>1<br>55<br>9<br>3                      | 12<br>31<br>35<br>35<br>35<br>70<br>40<br>50                 | 10<br>27<br>24<br>10<br>3<br>60<br>22<br>5                        | 12<br>31<br>35<br>35<br>32<br>67<br>40<br>50                  | 4.9%<br>9.5%<br>8.8%<br>10.5%<br>14.9%<br>14.5%<br>9.9%<br>15.4%                                      | -0.33   0.69, 0.02]<br>-0.45   0.66, -0.24]<br>-0.20   0.43, 0.03]<br>-0.14   -0.33, 0.05]<br>-0.07   -0.18, 0.05]<br>-0.11   -0.23, 0.01]<br>-0.33   -0.53, -0.12]<br>-0.04   -0.15, 0.07]              |      |  |    |
| Guan XG 2009<br>Jia JN 2018<br>Liu JG 2011<br>Wang L 2014<br>Wang Y 2017<br>Yao J 2017<br>Ye CY 2019<br>Zhang MM 2019<br>Total (95% Cl)  | 6<br>13<br>17<br>5<br>1<br>55<br>9<br>3                      | 12<br>31<br>35<br>35<br>35<br>70<br>40<br>50<br>339          | 10<br>27<br>24<br>10<br>3<br>60<br>22<br>5                        | 12<br>31<br>35<br>35<br>32<br>67<br>40<br>50<br>333           | 4.9%<br>9.5%<br>8.8%<br>10.5%<br>14.9%<br>14.5%<br>9.9%<br>15.4%                                      | -0.33 [0.69, 0.02]<br>-0.45 [0.66, 0.04]<br>-0.20 [0.43, 0.03]<br>-0.14 [0.33, 0.05]<br>-0.07 [0.18, 0.05]<br>-0.11 [-0.23, 0.01]<br>-0.33 [0.53, -0.12]<br>-0.04 [-0.15, 0.07]<br>-0.17 [-0.26, -0.08]  |      | •  |    |
| Guan XG 2009<br>Jia JN 2018<br>Liu JG 2011<br>Wang L 2014<br>Wang Y 2017<br>Yao J 2017<br>Yao Y 2019<br>Zhang MM 2019<br>Total (95% CI)<br>Total events                                      | 6<br>13<br>17<br>5<br>1<br>55<br>9<br>3                      | 12<br>31<br>35<br>35<br>35<br>70<br>40<br>50<br>339          | 10<br>27<br>24<br>10<br>3<br>60<br>22<br>5                        | 12<br>31<br>35<br>35<br>32<br>67<br>40<br>50<br>333           | 4.9%<br>9.5%<br>8.8%<br>10.5%<br>14.9%<br>14.5%<br>9.9%<br>15.4%                                      | -0.33 [0.68, 0.02]<br>-0.45 [0.66, 0.024]<br>-0.20 [0.43, 0.03]<br>-0.14 [0.33, 0.05]<br>-0.07 [0.18, 0.05]<br>-0.07 [0.18, 0.05]<br>-0.33 [-0.53, -0.12]<br>-0.04 [-0.15, 0.07]<br>-0.17 [-0.26, -0.08] |      | •  |    |
| Guan XG 2009<br>Jia JN 2018<br>Liu JG 2011<br>Wang L 2014<br>Wang Y 2017<br>Yao J 2017<br>Ye CY 2019<br>Zhang MM 2019<br>Total (95% CI)<br>Total events<br>Heterogeneity. Tau <sup>2</sup> = | 6<br>13<br>17<br>5<br>1<br>55<br>9<br>3<br>111<br>= 0.01; Ch | 12<br>31<br>35<br>35<br>35<br>70<br>40<br>50<br>339<br>#= 20 | 10<br>27<br>24<br>10<br>3<br>60<br>22<br>5<br>168<br>91. df = 8 ( | 12<br>31<br>35<br>35<br>32<br>67<br>40<br>50<br>333<br>P=0.00 | 4.9%<br>9.5%<br>8.8%<br>10.5%<br>14.9%<br>14.5%<br>9.8%<br>15.4%<br>100.0%<br>7); I <sup>2</sup> = 62 | -0.33 [0.69, 0.02]<br>-0.45 [0.66, 0.024]<br>-0.20 [0.43, 0.03]<br>-0.14 [0.33, 0.05]<br>-0.07 [0.18, 0.05]<br>-0.11 [0.23, 0.01]<br>-0.33 [-0.53, -0.12]<br>-0.04 [-0.15, 0.07]<br>-0.17 [-0.26, -0.08] | ļ    | •  | Ţ  |

Figure 4 The analysis of nausea and vomiting between two groups.









Table 3 Characteristics of studies with median survival time

| First author year  | NSCLC (III–IV) |       |         | MST           |             | 1-year su    | rvival rate  | ES (95% conf interval) |  |
|--------------------|----------------|-------|---------|---------------|-------------|--------------|--------------|------------------------|--|
| First aution, year | E/C            | M/F   | Age     | Age Treatment |             | Treatment    | Control      |                        |  |
| Guan 2009, (19)    | 12/12          | 11/12 | 36–72   | 18.1 months   | 14.3 months | Not reported | Not reported | 0.236 (-0.564, 1.036)  |  |
| Liu 2019, (28)     | 63/63          | 79/47 | 50–77   | 43.7 weeks    | 31.9 weeks  | 50.8%        | 34.9%        | 0.315 (-0.045, 0.674)  |  |
| Zhang 2019, (45)   | 50/50          | 52/48 | Unclear | 13.65 months  | 8.54 months | 48%          | 24%          | 0.469 (0.077, 0.861)   |  |

NSCLC, non-small cell lung cancer; MST, median survival time.



Figure 7 The analysis of KPS between two groups. KPS, Karnofsky performance status.

| Study of Subaroum  | Events  | Total  | Events  | Total  | Weight   | M-H. Fixed. 95% Cl   | M-H, Fixed, 95% C1  |
|--|---|--|---|--|--|--|---|
| Ban H 2019   | 30  | 31   | 25  | 31   | 2.9%   | 1 20 11 00 1 441   |   |
| Chen C 2018  | 24  | 30   | 20  | 30   | 2 394  | 1.20 (0.88 1.64)   | -   |
| Chan W 2016  | 27  | 44   | 27  | 44   | 31%  | 1 00 10 72 1 391   | +   |
| Chen V 2019  | 20  | 61   | 27  | 51   | 4 294  | 1 06 (0 84 1 22)   | +   |
| Chen 1 2010  | 28  | 20   | 31  | 31   | 4.370  | 1.05 [0.64, 1.32]  | 1   |
| Chen YZ 2003   | 22  | 28   | 23  | 21   | 2,7%   | 0.92 [0.72, 1.18]  | - I_  |
| Guan XQ 2009   | 11  | 12   | 8   | 12   | 0.9%   | 1.38 [0.89, 2.12]  |   |
| He LT 2017   | 48  | 54   | 33  | 54   | 3.8%   | 1.45 (1.15, 1.84)  |   |
| Huang ZB 2010  | 28  | 35   | 27  | 35   | 3.1%   | 1.04 [0.01, 1.32]  | +   |
| Jia JN 2018  | 24  | 31   | 21  | 31   | 2.4%   | 1.14 (0.84, 1.56)  | -   |
| LI HY 2017   | 25  | 41   | 25  | 41   | 2.9%   | 1.00 [0.71, 1.41]  | +   |
| LIL 2012   | 28  | 38   | 29  | 40   | 3.3%   | 1.02 (0.78, 1.33)  | -   |
| J XD 2016  | 34  | 39   | 25  | 39   | 2.9%   | 1.36 [1.04, 1.77]  |   |
| jang J 2018  | 36  | 40   | 31  | 40   | 3.6%   | 1 16 10 95 1 411   | +-  |
| iang SG 2014   | 18  | 23   | 15  | 20   | 1.9%   | 1 04 10 75 1 451   | +   |
| WE 2010  | 64  | 62   | 12  | 63   | 5 000  | 1 26 (1 02 1 52)   | -   |
| Ju 10 2015   | 29  | 25   | 20  | 25   | 2.0%   | 1.20 [1.00, 1.00]  | -   |
| 10.00 2011   | 40  | 35   | 20  | 30   | 5,0%   | 1.06 [0.65, 1.59]  | 1   |
| JUXH 2016  | 97  | 20   | 44  | 00   | 5.1%   | 1.07 [0.90, 1.27]  |   |
| LIU Y 2015   | 36  | 43   | 28  | 43   | 3.3%   | 1.29 (1.00, 1.66)  |   |
| .ong SG 2017   | 29  | 42   | 26  | 40   | 3.1%   | 1.06 [0.78, 1.44]  |   |
| du Q 2018  | 41  | 47   | 37  | 47   | 4.3%   | 1.11 [0.92, 1.33]  | +   |
| Sun SQ 2012  | 27  | 35   | 26  | 35   | 3.0%   | 1.04 [0.80, 1.35]  | +   |
| Nang HY 2016   | 13  | 24   | 12  | 25   | 1.4%   | 1.13 [0.65, 1.95]  |   |
| Nang L 2014  | 37  | 43   | 28  | 43   | 3.3%   | 1.32 [1.03. 1.70]  |   |
| Nang Y 2017  | 32  | 36   | 25  | 36   | 2.9%   | 1.32 11 04 1 671   | -   |
| (an QH 2019  | 20  | 40   | 20  | 40   | A 400  | 1.03 (0.02 1 36)   | +   |
| Vanal 2016   | 20  | 25   | 30  | 92   | 2 70   | 1 20 10 00 1 70  |   |
| ang E 2010   | 30  | 35   | 15  | 30   | 2.7%   | 1.30 [0.99, 1.72]  |   |
| ang 5J 2003  | 22  | 25   | 19  | 26   | 2.2%   | 1.20 [0.92, 1.58]  | L.  |
| rao J 2017   | 47  | 70   | 42  | 67   | 5.0%   | 1.07 [0.84, 1.37]  | T   |
| /e CY 2019   | 36  | 40   | 35  | 40   | 4,1%   | 1.03 [0.88, 1.20]  | T   |
| /u T 2015  | 52  | 60   | 38  | 60   | 4.4%   | 1.37 [1.10, 1.70]  |   |
| Zhang MM 2019  | 35  | 50   | 20  | 50   | 2.3%   | 1.75 [1.19, 2.57]  |   |
| otal (95% CI)  |   | 1249   |   | 1244   | 100.0%   | 1.16 [1.11, 1.22]  |   |
| Cotal events   | 1000  |  | 856   |  |  | 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1  |   |
| Telefogeneny. Chi"=  | 7-647   | - 30 (F  | 00011   | /0   |  |  | 0.01 0.1 1 10 10  |
| Fest for overall effect  | Z = 6.47  | (P < 0.0   | 0001)<br>Chemothe   | erapy  |  | Risk Difference  | 0.01 0.1 1 10 10<br>Favours (Kanglaite) Favours (Chemotherapy)<br>Risk Difference                       |
| feel for overall effect  | Z = 6.47<br>Kangla<br>Events  | (P < 0.0<br>aite<br>Total  | Chemothe  | erapy<br>Total   | Weight   | Risk Difference<br>M-H, Fixed, 95% Cl  | 0.01 0.1 1 10 10<br>Favours [Kanglaite] Favours [Chemotherapy]<br>Risk Difference<br>M-H, Fixed, 95% Cl |
| Test for overall effect<br>Study or Subgroup   | Z = 6.47<br>Kangla<br>Events<br>30  | (P < 0.0<br>aite<br><u>Total</u><br>31   | Chemothe<br>Events<br>25  | erapy<br>Total<br>31   | Weight<br>2.5%   | Risk Difference<br>M-H, Fixed, 95% CI<br>0,16 (0.01, 0.31)   | 0.01 0.1 1 10 10<br>Favours (Kanglaite) Favours (Chemotherapy)<br>Risk Difference<br>M-H, Fixed, 95% Cl |
| Test for overall effect<br>Study or Subgroup<br>Bao H 2019<br>Chen C 2018  | Z = 6.47<br>Kangla<br>Events<br>30<br>24  | = 30 (P<br>(P < 0.0<br>aite<br><u>Total</u><br>31<br>30  | Chemothe<br>Events<br>25<br>20  | erapy<br>Total<br>31<br>30   | Weight<br>2.5%<br>2.4%   | Risk Difference<br>M-H, Fixed, 95% CI<br>0.16 (0.01, 0.31)<br>0.13 (-0.09, 0.35)   | 0.01 0.1 1 10 10<br>Favours [Kanglaite] Favours [Chemotherapy]<br>Risk Difference<br>M.H. Fixed, 95% Cl |
| Test for overall effect<br>Study or Subgroup<br>Bao H 2019<br>Chen C 2018<br>Chen V 2016   | Z = 6.47<br>Kangla<br>Events<br>30<br>24<br>27  | = 30 (P<br>(P < 0.0<br><u>Total</u><br>31<br>30<br>44  | 25<br>20<br>27  | erapy<br>Total<br>31<br>30<br>44   | Weight<br>2.5%<br>2.4%<br>3.5%   | Risk Difference<br>M-H, Fixed, 95% Cl<br>0.16 (0.01, 0.31)<br>0.13 (-0.09, 0.35)<br>0.00 (-0.20, 0.20)   | 0.01 0.1 1 10 10<br>Favours [Kanglaite] Favours [Chemotherapy]<br>Risk Difference<br>M-H, Fixed, 95% Cl |
| Test for overall effect<br>Study or Subgroup<br>Bao H 2019<br>Chen C 2018<br>Chen W 2018<br>Chen V 2018  | Z = 6.47<br>Kangla<br>Events<br>30<br>24<br>27<br>30  | (P < 0.0<br>aite<br><u>Total</u><br>31<br>30<br>44<br>51   | 25<br>20<br>27<br>27  | erapy<br>Total<br>31<br>30<br>44<br>51   | Weight<br>2.5%<br>2.4%<br>3.5%<br>4.1%   | Risk Difference<br><u>M-H, Fixed, 95% Cl</u><br>0.16 (0.01, 0.31)<br>0.13 (-0.09, 0.35)<br>0.00 (-0.20, 0.20)<br>0.04 6.0 13, 0.21)  | 0.01 0.1 1 10 10<br>Favours [Kanglaite] Favours [Chemotherapy]<br>Risk Difference<br>M-H, Fixed, 95% Cl |
| Test for overall effect<br>Study or Subgroup<br>Bao H 2019<br>Chen C 2018<br>Chen V 2018<br>Chen V 2018<br>Chen V 2018   | Z = 6.47<br>Kangla<br>Events<br>30<br>24<br>27<br>39  | (P < 0.0<br>aite<br><u>Total</u><br>31<br>30<br>44<br>51<br>20   | 25<br>27<br>27<br>27<br>27<br>27<br>27  | 270<br>270<br>270<br>270<br>270<br>270<br>270<br>270<br>270<br>270   | Weight<br>2.5%<br>2.4%<br>3.5%<br>4.1%   | Risk Difference<br>M-H, Fixed, 95% CI<br>0.16 (0.01, 0.31)<br>0.13 (-0.09, 0.35)<br>0.00 (-0.20, 0.20)<br>0.04 (-0.13, 0.21)<br>0.07 (-0.27, 0.14)   | 0.01 0.1 1 10 10<br>Favours (Kanglaite) Favours (Chemotherapy)<br>Risk Difference<br>M.H. Fixed, 95% Cl |
| Test for overall effect<br>Study or Subaroup<br>Bao H 2019<br>Chen C 2018<br>Chen V 2018<br>Chen Y 2018<br>Chen Y 2018<br>Chen Y 2018  | Z = 6.47<br>Kangla<br>Events<br>30<br>24<br>27<br>39<br>22  | (P < 0.0<br>aite<br><u>Total</u><br>31<br>30<br>44<br>51<br>28   | 26,43,,1<br>00001)<br>Chemothe<br>Events<br>25<br>20<br>27<br>37<br>23  | 270<br>270<br>271<br>271<br>272<br>272<br>272<br>272<br>272<br>272<br>272<br>272   | Weight<br>2.5%<br>2.4%<br>3.5%<br>4.1%<br>2.2%   | Risk Difference<br>M-H, Fixed, 95% CI<br>0.16 [0.01, 0.31]<br>0.13 (-0.09, 0.35]<br>0.00 (-0.20, 0.20]<br>0.04 (-0.13, 0.21]<br>-0.07 (-0.27, 0.14]  | 0.01 0.1 1 10 10<br>Favours [Kanglaite] Favours [Chemotherapy]<br>Risk Difference<br>M-H, Fixed, 95% CI |
| Fest for overall effect<br>Study or Subgroup<br>3ao H 2019<br>Chen C 2018<br>Chen V 2016<br>Chen Y 2018<br>Chen Y 2018<br>Suan XO 2009   | Z = 6.47<br>Kangla<br>Events<br>30<br>24<br>27<br>39<br>22<br>11  | (P < 0.0<br>aite<br><u>Total</u><br>31<br>30<br>44<br>51<br>28<br>12   | 26,43,1<br>00001)<br>Chemothe<br>Events<br>25<br>20<br>27<br>37<br>23<br>8<br>8   | erapy<br>Total<br>31<br>30<br>44<br>51<br>27<br>12   | Weight<br>2.5%<br>2.4%<br>3.5%<br>4.1%<br>2.2%<br>1.0%   | Risk Difference<br>M-H. Fixed, 95% CI<br>0.16 [0.01, 0.31]<br>0.13 [-0.09, 0.35]<br>0.00 [-0.20, 0.23]<br>0.04 [-0.13, 0.21]<br>-0.07 [-0.27, 0.14]<br>0.25 [-0.06, 0.56]  | 0.01 0.1 1 10 10<br>Favours [Kanglaite] Favours [Chemotherapy]<br>Risk Difference<br>M-H, Fixed, 95% Cl |
| Fest for overall effect<br>Study or Subgroup<br>3ao H 2019<br>Chen C 2018<br>Chen V 2018<br>Chen V 2018<br>Chen V2013<br>Chen V2003<br>Juan XO 2009<br>He LT 2017  | Z = 6.47<br>Kangla<br>Events<br>30<br>24<br>27<br>39<br>22<br>11<br>48  | - 30 (P < 0.0<br>aite<br><u>Total</u><br>31<br>30<br>44<br>51<br>28<br>12<br>54  | 25<br>20<br>27<br>37<br>23<br>8<br>33   | erapy<br><u>Total</u><br>31<br>30<br>44<br>51<br>27<br>12<br>54  | Weight<br>2.5%<br>2.4%<br>3.5%<br>4.1%<br>2.2%<br>1.0%<br>4.3%   | Risk Difference<br>M-H, Fixed, 95% CI<br>0.13 (-0.09, 0.35)<br>0.00 (-0.20, 0.20)<br>0.04 (-0.13, 0.21)<br>-0.07 (-0.27, 0.14)<br>0.25 (-0.06, 0.58)<br>0.28 (0.12, 0.43)  | 0.01 0.1 1 10 10<br>Favours [Kanglaite] Favours [Chemotherapy]<br>Risk Difference<br>M.H. Fixed, 95% Cl |
| Study or Subgroup<br>Bao H 2019<br>Chen C 2018<br>Chen W 2018<br>Chen W 2018<br>Chen YZ 2013<br>Chen YZ 2003<br>Suan XC 2009<br>4 LT 2017<br>-Juang ZB 2010  | Z = 6.47<br>Kangla<br>Events<br>30<br>24<br>27<br>39<br>22<br>11<br>48<br>28  | (P < 0.0<br>(P < 0.0<br>31<br>30<br>44<br>51<br>28<br>12<br>54<br>35   | 25<br>20<br>27<br>37<br>23<br>8<br>33<br>27   | erapy<br>Total<br>31<br>30<br>44<br>51<br>27<br>12<br>54<br>35   | Weight<br>2.5%<br>2.4%<br>3.5%<br>4.1%<br>2.2%<br>1.0%<br>4.3%<br>2.8%   | Risk Difference<br>M-H, Fixed, 95% CI<br>0.16 [0.01, 0.31]<br>0.00 [-0.20, 0.20]<br>0.04 [-0.13, 0.21]<br>-0.07 [-0.27, 0.14]<br>0.25 [-0.06, 0.56]<br>0.28 [0.12, 0.43]<br>0.03 [-0.16, 0.22]   | 0.01 0.1 1 10 10<br>Favours [Kanglaite] Favours [Chemotherapy]<br>Risk Difference<br>M-H, Fixed, 95% CI |
| Feel of general effect<br>Study or Subproug<br>Jao H 2019<br>Chen C 2018<br>Chen V 2018<br>Chen Y 2018<br>Chen YZ 2003<br>Suan XO 2009<br>He LT 2017<br>Huang ZB 2010<br>Jia JN 2018   | Z = 6.47<br>Kangla<br>Events<br>30<br>24<br>27<br>39<br>22<br>11<br>48<br>28<br>24  | (P < 0.0<br>(P < 0.0<br>31<br>30<br>44<br>51<br>28<br>12<br>54<br>35<br>31   | 20,43), 100001)<br>Chemothe<br>Events<br>25<br>20<br>27<br>37<br>23<br>8<br>33<br>27<br>21  | 27 apy<br>Total<br>31<br>30<br>44<br>51<br>27<br>12<br>54<br>35<br>31  | Weight<br>2.5%<br>2.4%<br>3.5%<br>4.1%<br>2.2%<br>1.0%<br>4.3%<br>2.8%<br>2.5%   | Risk Difference<br>M-H, Fixed, 95% CI<br>0.16 [0.01, 0.31]<br>0.13 [-0.09, 0.35]<br>0.00 [-0.20, 0.20]<br>0.04 [-0.13, 0.21]<br>-0.07 [-0.27, 0.14]<br>0.25 [-0.06, 0.56]<br>0.28 [0.12, 0.43]<br>0.03 [-0.16, 0.22]<br>0.10 [-0.12, 0.32]   | 0.01 0.1 1 10 10<br>Favours [Kanglaite] Favours [Chemotherapy]<br>Risk Difference<br>M-H, Fixed, 95% Cl |
| First for overall effect<br>Study or Subgroup<br>Jao H 2019<br>Chen C 2018<br>Chen V 2018<br>Chen V 2018<br>Chen V 2013<br>Suan XO 2009<br>He LT 2017<br>Huang ZB 2010<br>Jia JN 2018<br>LI HY 2017  | Z = 6.47<br>Kangla<br>Events<br>30<br>24<br>27<br>39<br>22<br>11<br>11<br>48<br>28<br>24<br>25  | (P < 0.0<br>aite<br><u>Total</u><br>31<br>30<br>44<br>51<br>28<br>12<br>54<br>35<br>31<br>41   | 20001)<br>Chemothe<br>Events<br>25<br>20<br>27<br>37<br>23<br>8<br>33<br>27<br>21<br>25   | Total<br>31<br>30<br>44<br>51<br>27<br>12<br>54<br>35<br>31<br>41  | Weinht<br>2.5%<br>2.4%<br>3.5%<br>4.1%<br>2.2%<br>1.0%<br>4.3%<br>2.8%<br>2.8%<br>3.3%   | Risk Difference<br>M-H, Fixed, 95% CI<br>0.18 [0.01, 0.31]<br>0.13 [-0.09, 0.35]<br>0.00 [-0.20, 0.20]<br>0.04 [-0.13, 0.21]<br>-0.07 [-0.27, 0.14]<br>0.25 [-0.06, 0.56]<br>0.28 [0.12, 0.43]<br>0.03 [-0.16, 0.23]<br>0.10 [-0.12, 0.32]<br>0.00 [-0.21, 0.21]   | 0.01 0.1 1 10 10 Favours [Kanglaite] Favours [Chemotherapy] Risk Difference M.H. Fixed, 95% Cl          |
| Test for overall effect<br>Study or Suborous<br>3ao H 2019<br>Chen C 2018<br>Chen W 2018<br>Chen VZ 2003<br>Shen VZ 2003<br>Suan XO 2009<br>4 LT 2017<br>Huang ZB 2010<br>Ia JN 2018<br>LI HY 2017<br>JL 2012  | Z = 6.47<br>Kangla<br>Events<br>30<br>24<br>27<br>39<br>22<br>11<br>48<br>28<br>24<br>25<br>28  | (P < 0.0<br>aite<br>Total<br>31<br>30<br>44<br>51<br>28<br>12<br>54<br>35<br>31<br>41<br>38  | 20,433,1<br>00001)<br>Chemothe<br><u>Events</u><br>20<br>27<br>27<br>37<br>23<br>8<br>33<br>27<br>21<br>25<br>29  | erapy<br>Total<br>31<br>30<br>44<br>51<br>27<br>12<br>54<br>35<br>31<br>41<br>40   | Weight<br>2.5%<br>2.4%<br>3.5%<br>4.1%<br>2.2%<br>1.0%<br>4.3%<br>2.8%<br>2.5%<br>3.3%<br>3.1%   | Risk Difference<br>M-H, Fixed, 95% CI<br>0.16 [0.01, 0.31]<br>0.13 [-0.09, 0.35]<br>0.00 [-0.20, 0.20]<br>0.04 [-0.13, 0.21]<br>-0.07 [-0.27, 0.14]<br>0.25 [-0.06, 0.56]<br>0.28 [0.12, 0.43]<br>0.03 [-0.16, 0.22]<br>0.10 [-0.21, 0.21]<br>0.00 [-0.21, 0.21]   | 0.01 0.1 1 10 10<br>Favours [Kanglaite] Favours [Chemotherapy]<br>Risk Difference<br>M-H, Fixed, 95% Cl |
| Test for overall effect<br>Study or Subproug<br>3ao H 2019<br>Chen C 2018<br>Chen C 2018<br>Chen V 2018<br>Chen VZ 2003<br>3uan XO 2009<br>He LT 2017<br>Huang ZB 2010<br>Ha JN 2018<br>L HY 2017<br>LL 2012<br>JXD 2016   | Z = 6.47<br>Kangla<br>Events<br>30<br>24<br>27<br>38<br>22<br>11<br>48<br>28<br>24<br>25<br>28<br>24<br>25<br>28<br>34  | (P < 0.0<br>aite<br>Total<br>31<br>30<br>44<br>51<br>28<br>12<br>54<br>35<br>31<br>41<br>38<br>39  | Chemoth<br>Events<br>25<br>20<br>27<br>37<br>23<br>8<br>8<br>33<br>27<br>21<br>25<br>29<br>20<br>27<br>21<br>25<br>29<br>25   | Total<br>31<br>30<br>44<br>51<br>27<br>12<br>54<br>35<br>31<br>41<br>40<br>39  | Weialit<br>2.5%<br>2.4%<br>3.5%<br>4.1%<br>2.2%<br>4.3%<br>2.8%<br>2.8%<br>3.3%<br>3.1%  | Risk Difference<br>M-H, Fixed, 95% CI<br>0.16 [0.01, 0.31]<br>0.13 [-0.09, 0.35]<br>0.00 [-0.20, 0.20]<br>0.04 [-0.13, 0.21]<br>-0.07 [-0.27, 0.14]<br>0.25 [-0.06, 0.56]<br>0.28 [0.12, 0.43]<br>0.03 [-0.16, 0.22]<br>0.10 [-0.12, 0.32]<br>0.00 [-0.21, 0.21]<br>0.01 [-0.19, 0.21]<br>0.23 [0.05, 0.41]  | 0.01 0.1 1 10 10 Favours [Kanglaite] Favours [Chemotherapy] Risk Difference M.H. Fixed, 95% Cl          |
| Test for overall effect<br>Study or Subgroup<br>3ao H 2019<br>Chen C 2018<br>Chen V 2016<br>Chen Y 2018<br>Chen Y 2013<br>Suan XC 2009<br>He LT 2017<br>Huang ZB 2010<br>Ja JN 2018<br>J HY 2017<br>J L 2012<br>J XD 2018<br>Lang J 2018   | Z = 6.47<br>Kangla<br><u>Events</u><br>30<br>24<br>27<br>39<br>22<br>11<br>48<br>28<br>24<br>25<br>28<br>34<br>36   | (P < 0.0<br>aite<br>Total<br>31<br>30<br>44<br>51<br>28<br>12<br>54<br>35<br>31<br>41<br>35<br>31<br>41<br>39<br>40  | 25<br>20<br>27<br>27<br>37<br>23<br>8<br>33<br>27<br>21<br>25<br>29<br>20<br>27<br>37<br>23<br>37<br>23<br>8<br>33<br>27<br>21<br>25<br>29<br>25<br>25<br>31  | Total<br>31<br>30<br>44<br>51<br>27<br>12<br>54<br>35<br>31<br>41<br>40<br>39<br>40  | Weialitt<br>2.5%<br>2.4%<br>3.5%<br>4.1%<br>2.2%<br>1.0%<br>4.3%<br>2.8%<br>2.5%<br>3.3%<br>3.1%<br>3.1%<br>3.2%   | Risk Difference<br>M-H, Fixed, 95% CI<br>0.18 [0.01, 0.31]<br>0.13 {0.09, 0.35]<br>0.00 {-0.20, 0.20]<br>0.04 {-0.13, 0.21]<br>-0.07 {-0.27, 0.14]<br>0.25 {-0.06, 0.56]<br>0.28 [0.12, 0.43]<br>0.03 {-0.16, 0.22]<br>0.10 {-0.21, 0.32]<br>0.00 {-0.21, 0.21]<br>0.01 {-0.31, 0.31}<br>0.33 [0.05, 0.41]<br>0.13 {0.03, 0.28]  | 0.01 0.1 1 10 10 Favours [Kanglaite] Favours [Chemotherapy] Risk Difference M.H. Fixed, 95% Cl          |
| Study or Subgroup           Bao H 2019           Shen C 2018           Chen W 2016           Chen YZ 2003           Suan XO 2009           4 L T 2017           Huang ZB 2010           Ja J 2018           L HY 2017           L L 2017           Huang ZB 2010           Ja J 2018           Liang J 2018           Jang J 2018           Jang 2018           Jang 2018  | Z = 6.47<br>Kangla<br>Events<br>30<br>24<br>27<br>39<br>22<br>11<br>48<br>28<br>24<br>25<br>28<br>34<br>36<br>18  | (P < 0.0<br>aite<br>31<br>30<br>44<br>51<br>28<br>12<br>54<br>35<br>31<br>41<br>38<br>39<br>40<br>23   | Chemoth<br>Events<br>25<br>20<br>27<br>37<br>23<br>8<br>33<br>27<br>21<br>25<br>29<br>25<br>29<br>25<br>31<br>15  | Total<br>31<br>30<br>44<br>51<br>27<br>12<br>54<br>35<br>31<br>41<br>40<br>39<br>40<br>20  | Weight<br>2.5%<br>2.4%<br>3.5%<br>4.1%<br>2.2%<br>4.3%<br>2.8%<br>3.3%<br>3.1%<br>3.1%<br>3.1%<br>3.2%   | Risk Difference<br>M-H. Fixed, 95% CI<br>0.16 [0.01, 0.31]<br>0.00 [-0.20, 0.20]<br>0.04 [-0.13, 0.21]<br>-0.07 [-0.27, 0.14]<br>0.25 [-0.06, 0.56]<br>0.28 [0.12, 0.43]<br>0.03 [-0.16, 0.22]<br>0.10 [-0.21, 0.21]<br>0.00 [-0.21, 0.21]<br>0.23 [0.05, 0.41]<br>0.23 [0.05, 0.41]<br>0.3 [-0.22, 0.28]  | 0.01 0.1 1 10 10<br>Favours [Kanglaite] Favours [Chemotherapy]<br>Risk Difference<br>M-H, Fixed, 95% C1 |
| Test for overall effect<br>Study or Subgroup<br>3ao H 2019<br>Chen C 2018<br>Chen C 2018<br>Chen V 2018<br>Chen VZ 2003<br>3uan XO 2009<br>He LT 2017<br>Huang ZB 2010<br>Ha JN 2018<br>L HY 2017<br>LL 2012<br>LXD 2016<br>Lang J 2018<br>Lang J 2018<br>Lang SG 2014<br>Lu F 2019  | Z = 6.47<br>Kangla<br>Events<br>30<br>24<br>27<br>39<br>22<br>11<br>14<br>48<br>28<br>24<br>26<br>28<br>24<br>26<br>28<br>34<br>36<br>54<br>54<br>54<br>54<br>54<br>54<br>54<br>54<br>54<br>54  | (P < 0.0<br>aite<br>Total<br>31<br>30<br>44<br>51<br>28<br>54<br>35<br>31<br>41<br>38<br>39<br>40<br>283<br>83   | 00001)<br>Chemothe<br>Events<br>25<br>20<br>27<br>37<br>23<br>8<br>33<br>27<br>21<br>25<br>28<br>27<br>21<br>25<br>28<br>25<br>31<br>155<br>43  | Total<br>31<br>30<br>44<br>51<br>27<br>12<br>54<br>35<br>31<br>41<br>40<br>39<br>40<br>20<br>63  | Weight 2,5% 2,4% 3,5% 4,1% 2,2% 4,3% 2,8% 3,5% 3,1% 3,1% 3,1% 3,2% 1,7% 5,1%   | Risk Difference<br>M.H. Fixed, 95% CI<br>0.16 [0.01, 0.31]<br>0.13 [-0.09, 0.35]<br>0.00 [-0.20, 0.20]<br>0.04 [-0.13, 0.21]<br>-0.07 [-0.27, 0.14]<br>0.26 [-0.06, 0.56]<br>0.28 [0.12, 0.43]<br>0.03 [-0.16, 0.22]<br>0.10 [-0.12, 0.32]<br>0.00 [-0.21, 0.21]<br>0.21 [-0.19, 0.21]<br>0.23 [-0.05, 0.41]<br>0.13 [-0.03, 0.28]<br>0.03 [-0.22, 0.29]<br>0.17 [-0.03, 0.23]   | 0.01 0.1 1 10 10 Favours [Kanglaite] Favours [Chemotherapy] Risk Difference M.H. Fixed, 95% Cl          |
| Test for overall effect<br>Study or Subgroup<br>Bao H 2019<br>Shen C 2018<br>Shen V 2013<br>Shen Y 2013<br>Shen Y 2013<br>Shen Y 2013<br>Shen Y 2013<br>Shen Y 2013<br>Shen Y 2013<br>Jun XO 2009<br>4 LT 2017<br>Huang ZB 2010<br>Jun Y 2017<br>JL 2012<br>J XD 2018<br>Jang SG 2014<br>Jun F 2019<br>Jun Jun 2011  | Z = 6.47<br>Kents<br>30<br>24<br>27<br>39<br>22<br>11<br>48<br>24<br>25<br>28<br>34<br>36<br>18<br>34<br>24<br>25<br>28<br>34<br>36<br>18<br>34<br>24<br>26<br>28<br>34<br>36<br>36<br>36<br>36<br>36<br>36<br>37<br>39<br>30<br>30<br>30<br>30<br>30<br>30<br>30<br>30<br>30<br>30   | (P < 0.0<br>aite<br>Total<br>31<br>30<br>44<br>51<br>28<br>12<br>54<br>35<br>31<br>41<br>38<br>39<br>40<br>23<br>35<br>35  | Chemothe<br>Events<br>25<br>20<br>27<br>23<br>8<br>33<br>27<br>23<br>8<br>33<br>27<br>21<br>25<br>29<br>25<br>21<br>25<br>29<br>25<br>31<br>15<br>31<br>25  | erapy<br>Total<br>31<br>30<br>44<br>51<br>27<br>12<br>54<br>35<br>31<br>41<br>40<br>39<br>40<br>20<br>63<br>36   | Weight<br>2,5%<br>2,4%<br>3,5%<br>4,1%<br>2,2%<br>4,3%<br>2,5%<br>3,1%<br>3,1%<br>3,2%<br>1,7%<br>5,1%<br>2,8%   | Risk Difference<br>M-H, Fixed, 95% CI<br>0.18 [0.01, 0.31]<br>0.13 [-0.09, 0.35]<br>0.00 [-0.20, 0.20]<br>0.04 [-0.13, 0.21]<br>-0.07 [-0.27, 0.14]<br>0.25 [-0.06, 0.56]<br>0.28 [0.12, 0.43]<br>0.03 [-0.16, 0.22]<br>0.10 [-0.12, 0.32]<br>0.00 [-0.21, 0.21]<br>0.01 [-0.19, 0.21]<br>0.33 [-0.03, 0.28]<br>0.03 [-0.22, 0.29]<br>0.17 [0.30, 0.28]<br>0.03 [-0.22, 0.29]<br>0.17 [0.30, 0.3]<br>0.36 [-0.22, 0.29]<br>0.47 [-0.30, 0.28]<br>0.49 [-0.22, 0.29]<br>0.41 [-0.30, 0.28]<br>0.41 [-0.40, 0.30]<br>0.41 [-0.40, 0.30]<br>0.41 [-0.40, 0.30]<br>0.41 [-0.40, 0.30]<br>0.41 [-0.40, 0.30]<br>0.41 [-0.40, 0.40]<br>0.41 [-0.40,            | 0.01 0.1 1 10 10<br>Favours [Kanglaite] Favours [Chemotherapy]<br>Risk Difference<br>M-H, Fixed, 95% Cl |
| Pacebogenein, Chr =<br>est for overall effect<br>Study or Subgroup<br>Bao H 2019<br>Chen V 2018<br>Chen VZ 2003<br>Shen VZ 2003<br>Shen VZ 2003<br>Shen VZ 2003<br>Shen VZ 2003<br>Lang ZB 2010<br>I 2017<br>Huang ZB 2010<br>I 2012<br>J XD 2016<br>Lang J 2018<br>Jang SG 2014<br>Ju F 2019<br>Ju JQ 2011<br>IN XH 2015  | Z = 6.47<br>Kangla<br><u>Events</u><br>30<br>24<br>27<br>39<br>22<br>11<br>48<br>28<br>24<br>26<br>28<br>24<br>26<br>28<br>34<br>36<br>18<br>54<br>28<br>36<br>36<br>36<br>36<br>36<br>36<br>36<br>39<br>39<br>39<br>39<br>39<br>39<br>39<br>39<br>39<br>39   | (P < 0.0<br>aite<br><u>Total</u><br>31<br>30<br>44<br>51<br>28<br>12<br>54<br>35<br>31<br>41<br>38<br>39<br>40<br>23<br>83<br>35<br>55<br>54<br>55<br>54<br>55<br>54<br>55<br>55<br>55<br>5  | 0001)<br>Chemothu<br>Events<br>25<br>20<br>27<br>37<br>23<br>8<br>33<br>27<br>21<br>25<br>28<br>25<br>28<br>25<br>31<br>15<br>5<br>43<br>43<br>26<br>44   | erapy<br>Total<br>31<br>30<br>44<br>51<br>27<br>12<br>54<br>35<br>31<br>41<br>40<br>39<br>940<br>20<br>63<br>35<br>55  | Weinitt<br>2.5%<br>2.4%<br>3.5%<br>4.1%<br>2.2%<br>4.3%<br>2.8%<br>3.3%<br>3.1%<br>3.1%<br>3.2%<br>5.1%<br>5.1%<br>2.8%  | Risk Difference<br>M-H, Fixed, 95% CI<br>0.16 [0.01, 0.31]<br>0.13 [-0.09, 0.35]<br>0.00 [-0.20, 0.20]<br>0.04 [-0.13, 0.21]<br>-0.07 [-0.27, 0.14]<br>0.25 [-0.06, 0.56]<br>0.28 [0.12, 0.43]<br>0.03 [-0.16, 0.22]<br>0.10 [-0.12, 0.32]<br>0.00 [-0.21, 0.21]<br>0.23 [0.05, 0.41]<br>0.23 [0.05, 0.41]<br>0.13 [-0.03, 0.28]<br>0.03 [-0.22, 0.29]<br>0.17 [0.03, 0.32]<br>0.06 [-0.14, 0.25]<br>0.05 [-0.09, 0.27]<br>0.05 [-0.09,            | 0.01 0.1 1 10 10<br>Favours [Kanglaite] Favours [Chemotherapy]<br>Risk Difference<br>M-H, Fixed, 95% Cl |
| Test for overall effect<br>Study or Subgroup<br>Bao H 2019<br>Chen C 2018<br>Chen C 2018<br>Chen Y 2018<br>Chen YZ 2003<br>Buan XO 2009<br>He LT 2017<br>Uang ZB 2010<br>Ha JN 2018<br>L 4Y 2017<br>LL 2012<br>LXD 2018<br>Lang J 2018<br>Lang J 2018<br>Lang J 2018<br>Lang SG 2014<br>Lu F 2019<br>Lu JG 2011<br>Lu X2 2015  | Z = 6.47<br>Kangla<br>Events<br>30<br>24<br>27<br>39<br>22<br>11<br>48<br>24<br>28<br>24<br>25<br>28<br>34<br>36<br>18<br>54<br>47<br>28<br>34<br>36<br>18<br>54<br>47<br>28<br>28<br>28<br>29<br>29<br>29<br>29<br>29<br>29<br>29<br>29<br>29<br>29  | (P < 0.0<br>aite<br>Total<br>31<br>30<br>44<br>51<br>28<br>12<br>54<br>35<br>31<br>41<br>38<br>39<br>40<br>23<br>83<br>35<br>55  | 00001)<br>Chemothe<br>Events<br>25<br>20<br>27<br>27<br>23<br>8<br>33<br>27<br>21<br>25<br>20<br>27<br>23<br>8<br>33<br>27<br>21<br>25<br>28<br>25<br>20<br>27<br>37<br>23<br>8<br>33<br>27<br>21<br>25<br>20<br>27<br>23<br>8<br>33<br>27<br>21<br>25<br>20<br>27<br>23<br>8<br>33<br>27<br>21<br>25<br>20<br>27<br>27<br>23<br>8<br>33<br>27<br>21<br>25<br>20<br>27<br>27<br>23<br>8<br>33<br>27<br>21<br>25<br>20<br>27<br>27<br>23<br>8<br>33<br>27<br>21<br>25<br>26<br>27<br>27<br>27<br>21<br>25<br>26<br>27<br>27<br>27<br>27<br>27<br>27<br>27<br>27<br>27<br>27  | Total<br>31<br>30<br>44<br>51<br>27<br>12<br>54<br>31<br>41<br>40<br>39<br>40<br>20<br>63<br>35<br>55  | Weialit<br>2.5%<br>2.4%<br>3.5%<br>4.1%<br>2.2%<br>1.0%<br>4.3%<br>2.8%<br>3.3%<br>3.1%<br>3.2%<br>3.1%<br>5.1%<br>5.2%<br>4.4%  | Risk Difference<br>M-H, Fixed, 95% CI<br>0.16 [0.01, 0.31]<br>0.13 [-0.09, 0.35]<br>0.00 [-0.20, 0.20]<br>0.04 [-0.13, 0.21]<br>-0.07 [-0.27, 0.14]<br>0.26 [-0.06, 0.56]<br>0.28 [0.12, 0, 43]<br>0.03 [-0.16, 0.22]<br>0.10 [-0.12, 0.32]<br>0.00 [-0.21, 0.21]<br>0.21 [-0.19, 0.21]<br>0.23 [0.05, 0.41]<br>0.13 [-0.3, 0.28]<br>0.03 [-0.22, 0.29]<br>0.17 [-0.3, 0.32]<br>0.06 [-0.14, 0.25]<br>0.05 [-0.09, 0.20]   | 0.01 0.1 1 10 10 Favours [Kanglaite] Favours [Chemotherapy] Risk Difference M.H. Fixed, 95% Cl          |
| Study or Subgroup           Bao H 2019           Shen C 2018           Chen V 2013           Chen Y 2013           Shan XZ 2003           Suan XO 2009           4e LT 2017           Huang ZB 2010           Jia JN 2018           L HY 2017           JL 2012           J XD 2018           Lang JG 2018           Lang SG 2014           Ju F 2019           Ju XD 2011           Ju XD 2015           Ju XH 2015           Ju Y 2015   | Z = 6.47<br>Kangla<br>Events<br>30<br>24<br>27<br>39<br>22<br>11<br>46<br>28<br>24<br>27<br>39<br>22<br>24<br>27<br>39<br>22<br>24<br>27<br>39<br>22<br>24<br>27<br>39<br>22<br>24<br>27<br>39<br>22<br>24<br>27<br>39<br>22<br>24<br>27<br>39<br>22<br>24<br>27<br>39<br>22<br>24<br>27<br>39<br>22<br>24<br>27<br>39<br>22<br>24<br>27<br>39<br>22<br>24<br>27<br>39<br>22<br>24<br>27<br>39<br>22<br>24<br>27<br>39<br>22<br>24<br>27<br>39<br>22<br>24<br>26<br>28<br>28<br>24<br>28<br>24<br>28<br>24<br>28<br>28<br>28<br>28<br>28<br>28<br>28<br>28<br>28<br>28  | - 36 (), (P < 0, 0<br>aite<br>Total<br>31<br>30<br>44<br>451<br>28<br>12<br>54<br>54<br>31<br>31<br>38<br>39<br>9<br>40<br>23<br>31<br>39<br>40<br>23<br>35<br>55<br>55<br>43  | 0001)<br>Chemothi<br>Events<br>25<br>20<br>27<br>37<br>23<br>8<br>8<br>33<br>27<br>21<br>25<br>29<br>25<br>31<br>15<br>43<br>26<br>44<br>44<br>28   | erapy<br>Total<br>31<br>30<br>44<br>51<br>27<br>54<br>35<br>31<br>40<br>20<br>63<br>39<br>40<br>20<br>63<br>55<br>55<br>43   | Weialit<br>2.5%<br>2.4%<br>3.5%<br>4.1%<br>2.2%<br>1.0%<br>4.3%<br>2.8%<br>3.3%<br>3.1%<br>3.3%<br>3.1%<br>3.2%<br>5.1%<br>2.8%<br>3.5%  | Risk Difference<br>M-H, Fixed, 95% CI<br>0.18 [0.01, 0.31]<br>0.13 [-0.09, 0.35]<br>0.00 [-0.20, 0.20]<br>0.04 [-0.13, 0.21]<br>-0.07 [-0.27, 0.14]<br>0.25 [-0.06, 0.56]<br>0.28 [0.12, 0.43]<br>0.03 [-0.16, 0.22]<br>0.01 [-0.19, 0.32]<br>0.01 [-0.19, 0.32]<br>0.01 [-0.19, 0.21]<br>0.33 [-0.03, 0.28]<br>0.03 [-0.22, 0.29]<br>0.17 [0.03, 0.32]<br>0.05 [-0.09, 0.20]<br>0.19 [0.01, 0.37]<br>0.19 [0.01, 0.37]<br>0 | 0.01 0.1 1 10 10 Favours [Kanglaite] Favours [Chemotherapy] Risk Difference M.H. Fixed, 95% Ci          |
| Study or Subgroup           Bao H 2019           Shen C 2018           Chen W 2016           Chen YZ 2003           Suan XO 2009           Hang ZB 2010           Jao J 2017           Huang ZB 2010           Ja J 2018           LH Y 2017           LL 2017           Huang ZB 2010           Ja JN 2018           LHY 2017           LL 2012           JXD 2016           Lang J 2018           Jang SG 2014           Ju F 2019           Ju Y 2015           Jung Y 2015   | Z = 6.47<br>Kangla<br><u>Events</u><br>30<br>24<br>27<br>39<br>22<br>11<br>48<br>28<br>24<br>25<br>28<br>24<br>25<br>28<br>34<br>36<br>18<br>54<br>28<br>36<br>18<br>54<br>28<br>36<br>29<br>29<br>29<br>29<br>29<br>29<br>29<br>29<br>29<br>29   | - 36 (P < 0.0<br>aite<br>Total<br>31<br>31<br>31<br>30<br>30<br>44<br>451<br>12<br>28<br>54<br>54<br>35<br>54<br>31<br>38<br>39<br>90<br>23<br>35<br>55<br>43<br>42<br>24<br>23<br>55<br>55<br>43  | Chemothu<br>Events<br>25<br>20<br>27<br>37<br>23<br>8<br>33<br>27<br>21<br>25<br>20<br>27<br>21<br>25<br>20<br>27<br>21<br>25<br>20<br>27<br>21<br>25<br>20<br>27<br>21<br>25<br>20<br>27<br>21<br>23<br>8<br>33<br>27<br>21<br>25<br>20<br>27<br>21<br>25<br>20<br>27<br>27<br>23<br>8<br>20<br>27<br>21<br>25<br>20<br>27<br>27<br>23<br>26<br>20<br>27<br>27<br>23<br>26<br>20<br>27<br>27<br>23<br>26<br>26<br>20<br>27<br>27<br>23<br>26<br>26<br>20<br>27<br>27<br>23<br>26<br>26<br>26<br>26<br>27<br>27<br>26<br>26<br>26<br>26<br>27<br>27<br>26<br>26<br>26<br>26<br>26<br>27<br>26<br>26<br>26<br>26<br>26<br>26<br>26<br>26<br>26<br>26   | rapy<br><u>Total</u><br>31<br>30<br>44<br>51<br>27<br>12<br>24<br>35<br>31<br>41<br>40<br>20<br>63<br>35<br>55<br>43<br>40   | Weialitt<br>2.5%<br>2.4%<br>3.5%<br>4.1%<br>2.2%<br>1.0%<br>4.3%<br>2.8%<br>3.3%<br>3.1%<br>3.1%<br>3.1%<br>5.1%<br>2.8%<br>4.4%<br>3.3%<br>4.4%<br>3.5%<br>3.3%   | Risk Difference<br>M-H, Fixed, 95% CI<br>0.16 [0.01, 0.31]<br>0.13 [-0.09, 0.35]<br>0.00 [-0.20, 0.20]<br>0.04 [-0.13, 0.21]<br>-0.07 [-0.27, 0.14]<br>0.25 [-0.06, 0.56]<br>0.28 [0.12, 0.43]<br>0.03 [-0.16, 0.22]<br>0.10 [-0.12, 0.32]<br>0.00 [-0.21, 0.21]<br>0.23 [0.05, 0.41]<br>0.23 [0.05, 0.41]<br>0.32 [0.05, 0.41]<br>0.32 [0.02, 0.29]<br>0.17 [0.03, 0.32]<br>0.06 [-0.14, 0.25]<br>0.05 [-0.09, 0.20]<br>0.19 [0.01, 0.37]<br>0.44 [-0.16, 0.24]   | 0.01 0.1 1 10 10<br>Favours [Kanglaite] Favours [Chemotherapy]<br>Risk Difference<br>M-H, Fixed, 95% Cl |
| Techogoneny, Chr =<br>est for overall effect<br>Study or Subproup<br>and H 2019<br>Chen C 2018<br>Chen V 2018<br>Chen YZ 2003<br>Suan XO 2009<br>He LT 2017<br>Uang ZB 2010<br>Iia JN 2018<br>LAY 2017<br>LL 2012<br>LXD 2018<br>Lang J 2018<br>Lang J 2018<br>Lang SG 2014<br>Ju F 2019<br>Ju JG 2011<br>Ju YA 2015<br>Jung 2018<br>Lang SG 2017<br>Mu G 2018   | Z = 6.47<br>Kangla<br>Events<br>30<br>24<br>27<br>39<br>22<br>11<br>48<br>28<br>24<br>25<br>28<br>24<br>25<br>28<br>34<br>36<br>18<br>54<br>47<br>36<br>29<br>41<br>41<br>41<br>41<br>41<br>41<br>41<br>41<br>41<br>41  | - 30 (P < 0.0)<br>- 31 (P < 0.0)<br>- 31 (3)<br>- 31   | Chemothe<br>Events<br>25<br>20<br>27<br>37<br>23<br>8<br>33<br>37<br>21<br>25<br>29<br>25<br>31<br>15<br>29<br>25<br>31<br>15<br>43<br>26<br>44<br>28<br>26<br>44<br>28<br>37   | erapy<br>Total<br>31<br>30<br>44<br>51<br>27<br>12<br>54<br>35<br>54<br>35<br>31<br>41<br>40<br>39<br>9<br>40<br>200<br>63<br>35<br>55<br>54<br>34<br>0<br>40<br>40<br>40<br>40<br>40<br>40<br>40<br>40<br>40<br>40<br>40<br>40  | Weialit<br>2.5%<br>2.4%<br>3.5%<br>4.1%<br>2.2%<br>1.0%<br>4.3%<br>2.8%<br>3.3%<br>3.1%<br>3.2%<br>3.1%<br>5.2%<br>4.4%<br>3.5%<br>3.3%<br>3.8%  | Risk Difference<br>M-H, Fixed, 95% CI<br>0.16 [0.01, 0.31]<br>0.13 [-0.09, 0.35]<br>0.00 [-0.20, 0.20]<br>0.04 [-0.13, 0.21]<br>-0.07 [-0.27, 0.14]<br>0.25 [-0.06, 0.66]<br>0.28 [0.12, 0.43]<br>0.03 [-0.16, 0.22]<br>0.10 [-0.12, 0.32]<br>0.00 [-0.21, 0.32]<br>0.01 [-0.19, 0.21]<br>0.23 [0.05, 0.41]<br>0.13 [-0.03, 0.28]<br>0.03 [-0.22, 0.28]<br>0.05 [-0.09, 0.20]<br>0.19 [0.01, 0.37]<br>0.04 [-0.16, 0.24]<br>0.09 [-0.07, 0.24]   | 0.01 0.1 1 10 10 Favours [Kanglaite] Favours [Chemotherapy] Risk Difference M.H. Fixed, 95% Cl          |
| Performance of the second seco   | Z = 6.47<br>Kangla<br>Events<br>30<br>24<br>27<br>39<br>22<br>111<br>48<br>28<br>24<br>26<br>28<br>34<br>36<br>18<br>54<br>28<br>34<br>36<br>18<br>54<br>27<br>36<br>29<br>41<br>27<br>36<br>29<br>41<br>27<br>29<br>20<br>20<br>20<br>20<br>20<br>20<br>20<br>20<br>20<br>20   | - 36 (P < 0.0<br>aite<br><u>Total</u><br>31<br>30<br>44<br>51<br>128<br>28<br>35<br>54<br>35<br>54<br>35<br>31<br>41<br>38<br>39<br>40<br>23<br>35<br>55<br>43<br>35<br>55<br>43<br>42<br>47<br>35<br>55<br>43<br>42<br>47<br>47<br>47<br>47<br>47<br>47<br>47<br>47<br>47<br>47   | 0001)<br>Chemothu<br>Events<br>25<br>20<br>27<br>37<br>23<br>8<br>8<br>33<br>27<br>21<br>25<br>29<br>25<br>31<br>15<br>43<br>26<br>44<br>428<br>26<br>37<br>26  | Image: second system           Total           31           30           44           51           27           12           54           35           36           55           56           43           40           20           63           35           55           56           43           40           47           35   | Weialit 2.5% 2.4% 3.5% 4.1% 2.2% 1.0% 2.8% 3.3% 3.1% 3.1% 3.1% 5.1% 5.1% 2.8% 3.5% 3.3% 3.5% 3.3% 3.8% 3.8% 3.8% 3.8% 3.8% 3.8% 3.8  | Risk Difference<br>M-H, Fixed, 95% CI<br>0.16 [0.01, 0.31]<br>0.13 [-0.09, 0.35]<br>0.00 [-0.20, 0.20]<br>0.04 [-0.13, 0.21]<br>-0.07 [-0.27, 0.14]<br>0.25 [-0.06, 0.56]<br>0.28 [0.12, 0.43]<br>0.03 [-0.16, 0.22]<br>0.10 [-0.21, 0.21]<br>0.01 [-0.19, 0.21]<br>0.03 [-0.22, 0.29]<br>0.17 [0.03, 0.28]<br>0.03 [-0.22, 0.29]<br>0.17 [0.03, 0.22]<br>0.05 [-0.09, 0.20]<br>0.19 [0.01, 0.37]<br>0.04 [-0.17, 0.23]<br>0.06 [-0.17, 0.23]  | 0.01 0.1 1 10 10 Favours [Kanglaite] Favours [Chemotherapy] Risk Difference M.H. Fixed, 95% Ci          |
| Participantial of the second s   | Z = 6.47<br>Kangla<br><u>Events</u><br>30<br>24<br>27<br>39<br>22<br>11<br>48<br>28<br>24<br>25<br>28<br>24<br>25<br>28<br>34<br>36<br>18<br>54<br>28<br>34<br>36<br>18<br>54<br>28<br>36<br>29<br>47<br>36<br>27<br>27<br>39<br>27<br>27<br>27<br>27<br>27<br>27<br>27<br>27<br>27<br>27   | - 30 (P < 0.0<br>(P < 0.0<br>31<br>30<br>44<br>51<br>32<br>54<br>35<br>54<br>31<br>31<br>32<br>54<br>31<br>32<br>54<br>31<br>32<br>54<br>31<br>32<br>54<br>31<br>32<br>54<br>31<br>31<br>30<br>44<br>41<br>31<br>30<br>44<br>42<br>54<br>43<br>55<br>54<br>43<br>44<br>41<br>51<br>54<br>54<br>54<br>54<br>54<br>54<br>54<br>54<br>54<br>54  | Chemothin<br>Events<br>25<br>20<br>27<br>27<br>37<br>23<br>8<br>33<br>27<br>21<br>25<br>25<br>31<br>33<br>27<br>21<br>25<br>25<br>31<br>15<br>15<br>43<br>26<br>44<br>44<br>28<br>26<br>37<br>26<br>43<br>26<br>43<br>26<br>43<br>26<br>43<br>26<br>43<br>26<br>43<br>26<br>43<br>26<br>25<br>20<br>27<br>77<br>77<br>23<br>8<br>8<br>33<br>27<br>27<br>27<br>27<br>27<br>27<br>27<br>27<br>27<br>27<br>27<br>27<br>27  | <b>Total</b><br>31<br>30<br>44<br>51<br>27<br>12<br>54<br>54<br>35<br>31<br>41<br>40<br>20<br>63<br>355<br>55<br>43<br>40<br>47<br>7<br>355<br>25<br>25  | Weialitt<br>2.5%<br>2.4%<br>3.5%<br>4.1%<br>2.2%<br>1.0%<br>4.3%<br>2.8%<br>3.3%<br>3.1%<br>3.1%<br>3.2%<br>1.7%<br>2.8%<br>4.4%<br>3.5%<br>3.3%<br>3.8%<br>3.8%<br>2.8%<br>2.2%   | Risk Difference<br>M-H, Fixed, 95% CI<br>0.16 [0.01, 0.31]<br>0.13 [-0.09, 0.35]<br>0.00 [-0.20, 0.20]<br>0.04 [-0.13, 0.21]<br>-0.07 [-0.27, 0.14]<br>0.25 [-0.06, 0.56]<br>0.28 [0.12, 0.43]<br>0.03 [-0.16, 0.22]<br>0.10 [-0.21, 0.21]<br>0.23 [0.05, 0.41]<br>0.23 [0.05, 0.41]<br>0.23 [0.05, 0.41]<br>0.32 [0.05, 0.41]<br>0.32 [0.05, 0.41]<br>0.32 [0.02, 0.29]<br>0.17 [0.03, 0.32]<br>0.06 [-0.14, 0.25]<br>0.05 [-0.09, 0.20]<br>0.19 [-0.07, 0.24]<br>0.03 [-0.17, 0.23]<br>0.04 [-0.16, 0.24]<br>0.03 [-0.17, 0.23]<br>0.06 [-0.22, 0.34]  | 0.01 0.1 1 10 10 Favours [Kanglaite] Favours [Chemotherapy] Risk Difference M.H. Fixed, 95% Cl          |
| Techogoneny, Chr =<br>Test for overall effect<br>Study or Subgroup<br>Sao H 2019<br>Chen C 2018<br>Chen V 2018<br>Chen YZ 2003<br>Suan XO 2009<br>He LT 2017<br>Uang ZE 2010<br>Ha Z018<br>L 2012<br>L 2013<br>L 2014<br>L 2015<br>Long SG 2017<br>Mu G 2018<br>Sun SQ 2012<br>Vang HZ 2016<br>Vang L 2014<br>Vang L 2016<br>Vang L 2014   | Z = 6.47<br>Kangla<br>Events<br>Events<br>27<br>30<br>24<br>27<br>39<br>22<br>111<br>48<br>24<br>27<br>38<br>24<br>27<br>39<br>22<br>111<br>48<br>24<br>27<br>38<br>24<br>27<br>39<br>22<br>111<br>48<br>24<br>27<br>39<br>22<br>111<br>48<br>24<br>25<br>28<br>34<br>36<br>29<br>21<br>111<br>48<br>24<br>25<br>28<br>34<br>36<br>29<br>21<br>111<br>48<br>24<br>25<br>28<br>34<br>36<br>29<br>29<br>21<br>111<br>48<br>24<br>25<br>28<br>34<br>36<br>26<br>27<br>27<br>28<br>28<br>26<br>28<br>34<br>26<br>27<br>27<br>28<br>28<br>24<br>26<br>28<br>34<br>36<br>26<br>27<br>28<br>28<br>34<br>36<br>26<br>27<br>27<br>28<br>28<br>34<br>36<br>26<br>27<br>27<br>28<br>34<br>36<br>26<br>27<br>28<br>34<br>36<br>26<br>27<br>27<br>28<br>34<br>36<br>29<br>27<br>28<br>34<br>36<br>29<br>29<br>29<br>29<br>29<br>20<br>20<br>20<br>20<br>20<br>20<br>20<br>20<br>20<br>20   | - 30 (P < 0.0<br>(P < 0.0<br>31<br>30<br>30<br>44<br>45<br>12<br>28<br>41<br>28<br>45<br>45<br>41<br>38<br>39<br>40<br>23<br>35<br>55<br>55<br>55<br>24<br>42<br>47<br>35<br>24<br>43<br>35<br>55<br>24<br>43<br>55<br>24<br>43<br>55<br>55<br>24<br>44<br>43<br>35<br>55<br>55<br>24<br>44<br>43<br>55<br>55<br>55<br>24<br>44<br>44<br>55<br>55<br>55<br>55<br>24<br>44<br>45<br>55<br>55<br>55<br>55<br>24<br>45<br>55<br>55<br>55<br>55<br>55<br>55<br>55<br>55<br>5   | Chemothin<br>Events<br>25<br>20<br>27<br>37<br>23<br>8<br>33<br>37<br>23<br>8<br>33<br>37<br>21<br>25<br>29<br>25<br>31<br>15<br>29<br>25<br>31<br>15<br>43<br>26<br>44<br>28<br>26<br>44<br>28<br>26<br>37<br>26<br>27<br>21<br>25<br>29<br>25<br>20<br>27<br>27<br>21<br>27<br>20<br>27<br>27<br>23<br>27<br>23<br>27<br>20<br>27<br>27<br>27<br>27<br>27<br>27<br>27<br>27<br>27<br>27<br>27<br>27<br>27   | Iterapy           Total           31           30           44           51           27           12           54           35           31           40           203           39           40           200           355           555           43           40           47           355           255           43           443  | Weight 2,5% 2,4% 3,5% 4,1% 2,2% 4,3% 2,2% 4,3% 2,8% 3,1% 3,1% 3,1% 3,1% 3,1% 5,1% 2,8% 4,4% 3,5% 3,3% 2,8% 2,8% 2,8% 2,5% 3,5%   | Risk Offference<br>M-H, Fixed, 95% CI<br>0.18 [0.01, 0.35]<br>0.00 [-0.20, 0.20]<br>0.04 [-0.13, 0.21]<br>-0.07 [-0.27, 0.14]<br>0.25 [-0.06, 0.56]<br>0.28 [0.12, 0.43]<br>0.03 [-0.16, 0.22]<br>0.10 [-0.12, 0.32]<br>0.00 [-0.21, 0.21]<br>0.01 [-0.19, 0.21]<br>0.03 [-0.22, 0.29]<br>0.17 [0.03, 0.32]<br>0.05 [-0.09, 0.20]<br>0.19 [0.01, 0.37]<br>0.04 [-0.16, 0.24]<br>0.04 [-0.16, 0.24]<br>0.04 [-0.17, 0.23]<br>0.06 [-0.12, 0.29]<br>0.04 [-0.16, 0.24]<br>0.04 [-0.17, 0.23]<br>0.06 [-0.22, 0.34]<br>0.06 [-0.22, 0.34]<br>0.05 [-0.03, 0.39]   | 0.01 0.1 1 10 10 Favours [Kanglaite] Favours [Chemotherapy] Risk Difference M.H. Fixed, 95% Cl          |
| Performance of the second seco   | Z = 6.47<br>Kangla<br>Events<br>30<br>24<br>27<br>39<br>22<br>21<br>11<br>48<br>28<br>24<br>26<br>28<br>34<br>36<br>18<br>54<br>28<br>34<br>36<br>29<br>41<br>127<br>13<br>37<br>33<br>37<br>33   | - 30 (P < 0.0<br>(P < 0.0<br>31<br>30<br>31<br>30<br>44<br>451<br>28<br>12<br>54<br>35<br>55<br>55<br>55<br>55<br>55<br>43<br>35<br>55<br>55<br>55<br>43<br>42<br>47<br>35<br>24<br>43<br>63<br>43<br>43<br>43<br>55<br>55<br>55<br>55<br>55<br>55<br>55<br>55<br>55<br>5  | Chemothin<br>Events<br>25<br>20<br>27<br>37<br>23<br>8<br>33<br>27<br>21<br>25<br>29<br>25<br>31<br>15<br>43<br>26<br>44<br>428<br>26<br>37<br>26<br>12<br>28<br>26<br>27<br>21<br>29<br>25<br>31<br>31<br>29<br>25<br>31<br>29<br>25<br>31<br>29<br>25<br>29<br>25<br>20<br>27<br>27<br>23<br>33<br>27<br>21<br>29<br>25<br>20<br>27<br>27<br>23<br>33<br>27<br>21<br>25<br>20<br>27<br>27<br>23<br>23<br>27<br>23<br>28<br>20<br>27<br>27<br>23<br>29<br>27<br>20<br>27<br>27<br>23<br>29<br>20<br>27<br>27<br>23<br>29<br>29<br>20<br>27<br>27<br>23<br>29<br>29<br>20<br>27<br>27<br>23<br>29<br>20<br>27<br>27<br>23<br>29<br>29<br>29<br>29<br>20<br>27<br>27<br>29<br>29<br>29<br>29<br>29<br>29<br>29<br>29<br>29<br>29<br>29<br>29<br>29   | Prapy           Total           31           300           44           51           27           12           24           355           355           355           355           355           355           355           355           355           355           355           355           355           355           355           355           355           355           356           357           358           359           350           351           355           355           355           356           357           358           358           358           359           350           350           351           352           356           357           358           359           350 <t< td=""><td>Weialit 2,5% 2,4% 3,5% 4,1% 2,2% 1,0% 2,2% 3,3% 3,1% 3,2% 3,1% 3,1% 3,1% 3,1% 3,1% 3,1% 3,1% 3,2% 3,3% 3,3% 3,3% 3,2% 3,2% 3,2% 3,2</td><td>Risk Difference<br/>M-H, Fixed, 95% CI<br/>0.16 [0.01, 0.31]<br/>0.13 [-0.09, 0.36]<br/>0.00 [-0.20, 0.20]<br/>0.04 [-0.13, 0.21]<br/>-0.07 [-0.27, 0.14]<br/>0.25 [-0.06, 0.56]<br/>0.28 [0.12, 0.43]<br/>0.03 [-0.16, 0.22]<br/>0.10 [-0.21, 0.32]<br/>0.01 [-0.19, 0.32]<br/>0.01 [-0.19, 0.21]<br/>0.33 [-0.32, 0.32]<br/>0.04 [-0.21, 0.21]<br/>0.33 [-0.32, 0.32]<br/>0.05 [-0.03, 0.32]<br/>0.05 [-0.09, 0.20]<br/>0.19 [0.01, 0.37]<br/>0.04 [-0.17, 0.23]<br/>0.06 [-0.22, 0.34]<br/>0.03 [-0.17, 0.23]<br/>0.06 [-0.22, 0.34]<br/>0.21 [0.03, 0.39]<br/>0.22 [0.05, 0.40]</td><td>0.01 0.1 1 10 10 Favours [Kanglaite] Favours [Chemotherapy] Risk Difference M.H. Fixed, 95% Ci</td></t<>   | Weialit 2,5% 2,4% 3,5% 4,1% 2,2% 1,0% 2,2% 3,3% 3,1% 3,2% 3,1% 3,1% 3,1% 3,1% 3,1% 3,1% 3,1% 3,2% 3,3% 3,3% 3,3% 3,2% 3,2% 3,2% 3,2  | Risk Difference<br>M-H, Fixed, 95% CI<br>0.16 [0.01, 0.31]<br>0.13 [-0.09, 0.36]<br>0.00 [-0.20, 0.20]<br>0.04 [-0.13, 0.21]<br>-0.07 [-0.27, 0.14]<br>0.25 [-0.06, 0.56]<br>0.28 [0.12, 0.43]<br>0.03 [-0.16, 0.22]<br>0.10 [-0.21, 0.32]<br>0.01 [-0.19, 0.32]<br>0.01 [-0.19, 0.21]<br>0.33 [-0.32, 0.32]<br>0.04 [-0.21, 0.21]<br>0.33 [-0.32, 0.32]<br>0.05 [-0.03, 0.32]<br>0.05 [-0.09, 0.20]<br>0.19 [0.01, 0.37]<br>0.04 [-0.17, 0.23]<br>0.06 [-0.22, 0.34]<br>0.03 [-0.17, 0.23]<br>0.06 [-0.22, 0.34]<br>0.21 [0.03, 0.39]<br>0.22 [0.05, 0.40]  | 0.01 0.1 1 10 10 Favours [Kanglaite] Favours [Chemotherapy] Risk Difference M.H. Fixed, 95% Ci          |
| Participantial, Chr =<br>Pest for overall effect<br>Study or Subgroup<br>Bao H 2019<br>Chen C 2018<br>Chen YZ 2003<br>Shen YZ 2003<br>Suan XG 2009<br>He LT 2017<br>LI 2017<br>LI 2017<br>LI 2012<br>LI 2012<br>LI 2012<br>LI 2012<br>LI 2012<br>LI 2012<br>LI 2013<br>Li 2013<br>Li 2013<br>Li 2014<br>JU F 2019<br>Li 4 2018<br>Sun SQ 2012<br>Vang YZ 2016<br>Vang YZ 2016<br>Vang L 2014<br>Vang YZ 2017<br>Yan QH 2018  | Z = 6.47<br>Kangla<br><u>Events</u><br>30<br>24<br>27<br>39<br>22<br>11<br>48<br>28<br>24<br>25<br>28<br>24<br>25<br>28<br>34<br>36<br>18<br>54<br>28<br>34<br>36<br>18<br>54<br>29<br>47<br>36<br>29<br>47<br>30<br>27<br>39<br>22<br>11<br>48<br>28<br>28<br>24<br>25<br>28<br>34<br>36<br>28<br>28<br>28<br>28<br>28<br>28<br>28<br>28<br>28<br>28   | - 30 (P < 0.0<br>(P < 0.0<br>31<br>30<br>30<br>44<br>51<br>28<br>31<br>28<br>31<br>28<br>35<br>55<br>55<br>55<br>55<br>55<br>55<br>55<br>55<br>55  | Chemothin<br>Events<br>25<br>20<br>27<br>27<br>23<br>8<br>33<br>27<br>21<br>25<br>20<br>27<br>23<br>8<br>33<br>27<br>21<br>25<br>25<br>31<br>15<br>43<br>26<br>44<br>26<br>43<br>26<br>43<br>26<br>43<br>26<br>43<br>26<br>43<br>26<br>43<br>26<br>43<br>26<br>43<br>26<br>43<br>26<br>43<br>26<br>43<br>26<br>43<br>26<br>43<br>26<br>43<br>26<br>43<br>26<br>43<br>26<br>43<br>26<br>43<br>26<br>43<br>26<br>43<br>26<br>43<br>26<br>27<br>27<br>21<br>25<br>26<br>27<br>27<br>27<br>27<br>27<br>27<br>27<br>27<br>27<br>27   | rapy<br>Total<br>31<br>30<br>44<br>451<br>27<br>12<br>27<br>12<br>27<br>12<br>27<br>12<br>35<br>31<br>41<br>40<br>39<br>40<br>20<br>63<br>35<br>55<br>55<br>55<br>55<br>55<br>55<br>43<br>40<br>40<br>40<br>40<br>40<br>40<br>40<br>40<br>40<br>40   | Weialitt<br>2.5%<br>2.4%<br>3.5%<br>4.1%<br>2.2%<br>4.3%<br>2.5%<br>3.3%<br>3.1%<br>3.1%<br>3.2%<br>4.4%<br>3.5%<br>3.3%<br>3.8%<br>2.8%<br>3.3%<br>3.8%<br>2.8%<br>3.5%<br>3.9%<br>3.9%   | Risk Difference<br>M-H, Fixed, 95% CI<br>0.16 [0.01, 0.31]<br>0.13 [-0.09, 0.35]<br>0.00 [-0.20, 0.20]<br>0.04 [-0.13, 0.21]<br>-0.07 [-0.27, 0.14]<br>0.25 [-0.06, 0.56]<br>0.28 [0.12, 0.43]<br>0.03 [-0.16, 0.22]<br>0.10 [-0.12, 0.32]<br>0.00 [-0.21, 0.21]<br>0.23 [0.05, 0.41]<br>0.13 [-0.13, 0.28]<br>0.03 [-0.22, 0.29]<br>0.17 [0.03, 0.32]<br>0.06 [-0.14, 0.25]<br>0.05 [-0.09, 0.20]<br>0.19 [-0.07, 0.24]<br>0.03 [-0.27, 0.24]<br>0.04 [-0.16, 0.24]<br>0.03 [-0.27, 0.24]<br>0.05 [-0.20, 0.30]<br>0.06 [-0.22, 0.34]<br>0.06 [-0.22, 0.34]<br>0.21 [0.03, 0.39]<br>0.22 [0.05, 0.40]   | 0.01 0.1 1 10 10 Favours [Kanglaite] Favours [Chemotherapy] Risk Difference M.H. Fixed, 95% Cl          |
| Test for overall effect<br>Study or Subgroup<br>Bao H 2019<br>Shen C 2018<br>Shen V 2013<br>Shen V 2013<br>Shen Y 2013<br>Shen Y 2013<br>Shen Y 2013<br>Shen Y 2013<br>Shen Y 2013<br>Shen Y 2017<br>Huang ZB 2010<br>HY 2017<br>H 2017<br>JL 2012<br>JXD 2018<br>Jang SG 2014<br>JJ Q 2015<br>JU XD 2014<br>Yang H 2016<br>Yang Y 2017<br>Yang H 2016<br>Yang Y 2017<br>Yang H 2018   | Z = 6.47<br>Kangla<br>Events<br>24<br>27<br>39<br>22<br>111<br>48<br>28<br>24<br>26<br>28<br>34<br>36<br>28<br>34<br>36<br>29<br>41<br>17<br>36<br>29<br>41<br>17<br>33<br>33<br>39<br>30<br>30<br>30<br>30<br>30<br>30<br>30<br>30<br>30<br>30   | Solution           Total           31           300           44           51           28           122           54           35           34           35           34           35           34           35           34           35           34           36           47           35           24           36           49           35   | Chemothic<br>Events<br>25<br>20<br>27<br>37<br>23<br>8<br>8<br>33<br>27<br>21<br>25<br>29<br>25<br>29<br>25<br>29<br>25<br>31<br>15<br>43<br>26<br>44<br>428<br>26<br>44<br>28<br>26<br>12<br>28<br>25<br>37<br>25<br>29<br>25<br>29<br>25<br>29<br>25<br>29<br>25<br>29<br>25<br>29<br>25<br>29<br>25<br>29<br>25<br>20<br>27<br>27<br>23<br>23<br>25<br>29<br>25<br>20<br>27<br>27<br>23<br>20<br>27<br>23<br>23<br>25<br>20<br>27<br>27<br>23<br>23<br>25<br>20<br>20<br>27<br>27<br>23<br>23<br>25<br>20<br>20<br>27<br>25<br>20<br>20<br>27<br>25<br>20<br>20<br>27<br>25<br>20<br>20<br>27<br>25<br>20<br>20<br>27<br>25<br>20<br>20<br>27<br>25<br>20<br>20<br>27<br>25<br>20<br>20<br>27<br>25<br>20<br>20<br>25<br>20<br>25<br>20<br>25<br>20<br>25<br>20<br>25<br>20<br>25<br>20<br>25<br>20<br>25<br>20<br>25<br>20<br>25<br>20<br>25<br>20<br>25<br>20<br>25<br>20<br>25<br>20<br>25<br>20<br>25<br>20<br>25<br>25<br>20<br>25<br>25<br>20<br>25<br>25<br>20<br>25<br>25<br>20<br>25<br>25<br>20<br>25<br>25<br>20<br>25<br>25<br>25<br>20<br>25<br>25<br>25<br>25<br>25<br>25<br>25<br>25<br>25<br>25<br>25<br>25<br>25  | rapy<br>Total<br>31<br>300<br>44<br>51<br>27<br>12<br>54<br>35<br>54<br>35<br>31<br>40<br>200<br>63<br>35<br>55<br>55<br>55<br>55<br>55<br>43<br>45<br>43<br>45<br>43<br>45<br>44<br>40<br>40<br>40<br>40<br>40<br>40<br>40<br>40<br>40  | Weight 2,5% 2,4% 3,5% 4,1% 2,2% 4,3% 2,2% 4,3% 2,8% 3,1% 3,1% 3,2% 3,1% 3,1% 3,2% 3,5% 3,3% 3,5% 3,3% 3,5% 2,8% 2,0% 3,2% 2,8% 2,9% 3,2% 2,9% 3,2% 2,2%  | Risk Difference<br>M-H, Fixed, 95% CI<br>0.18 [0.01, 0.35]<br>0.00 [-0.20, 0.20]<br>0.04 [-0.13, 0.21]<br>-0.07 [-0.27, 0.14]<br>0.25 [-0.06, 0.56]<br>0.28 [0.12, 0.43]<br>0.03 [-0.16, 0.22]<br>0.10 [-0.12, 0.32]<br>0.00 [-0.21, 0.21]<br>0.01 [-0.13, 0.32]<br>0.03 [-0.22, 0.29]<br>0.17 [0.03, 0.32]<br>0.05 [-0.09, 0.20]<br>0.19 [0.01, 0.37]<br>0.04 [-0.16, 0.24]<br>0.05 [-0.09, 0.20]<br>0.19 [0.01, 0.37]<br>0.04 [-0.17, 0.23]<br>0.06 [-0.22, 0.24]<br>0.03 [-0.17, 0.23]<br>0.06 [-0.22, 0.34]<br>0.03 [-0.17, 0.23]<br>0.06 [-0.22, 0.34]<br>0.22 [0.05, 0.40]<br>0.22 [0.05, 0.40]<br>0.22 [0.05, 0.40]<br>0.22 [0.00, 0.40]<br>0.22 [0.00, 0.40]<br>0.20 [0.00, 0.   | 0.01 0.1 1 10 10 Favours [Kanglaite] Favours [Chemotherapy] Risk Difference M.H. Fixed, 95% Ci          |
| Test for overall effect<br>Study or Suboroug<br>Bao H 2019<br>Chen C 2018<br>Chen W 2018<br>Chen V 2018<br>Chen V 2018<br>Chen VZ 2003<br>Buan XG 2009<br>4 LT 2017<br>Huang ZB 2010<br>Ja JN 2018<br>Jang G 2014<br>Ju F 2019<br>Ju JQ 2011<br>Ju Z 2018<br>Jun XH 2016<br>Jun Y 2015<br>Jun XH 2016<br>Jun Y 2015<br>Jun SQ 2012<br>Nang L 2018<br>Sun SQ 2012<br>Nang L 2018<br>Sun SQ 2012<br>Nang L 2018<br>Sun SQ 2012<br>Nang L 2014<br>Nang Y 2017<br>(an QH 2018<br>(an QH 2018<br>Cong SG 2017<br>(an QH 2018<br>Sun SQ 2012<br>Nang L 2017<br>(an QH 2018<br>Sun SQ 2017<br>(an QH 2018<br>(an QH 2018<br>(an QH 2018)<br>(an QH 2018<br>(an QH 2018)<br>(an QH 2018<br>(an QH 2018)<br>(an Q       | Z = 6.47<br>Kangla<br><u>Events</u><br>30<br>24<br>30<br>24<br>27<br>39<br>22<br>111<br>48<br>28<br>24<br>26<br>36<br>18<br>54<br>36<br>18<br>54<br>36<br>18<br>54<br>36<br>29<br>47<br>36<br>29<br>47<br>36<br>29<br>36<br>37<br>36<br>36<br>36<br>36<br>36<br>36<br>36<br>36<br>36<br>36  | - 30 (0 × 0.0.<br><b>Total</b><br>31<br>300<br>44<br>35<br>328<br>329<br>40<br>339<br>40<br>34<br>328<br>339<br>40<br>34<br>34<br>35<br>55<br>55<br>55<br>24<br>43<br>35<br>55<br>55<br>55<br>55<br>55<br>55<br>55<br>55<br>5  | Chemothin<br>Events<br>25<br>20<br>27<br>37<br>23<br>8<br>33<br>27<br>23<br>8<br>33<br>27<br>21<br>25<br>29<br>25<br>31<br>15<br>43<br>26<br>44<br>428<br>26<br>37<br>26<br>15<br>43<br>26<br>44<br>428<br>26<br>37<br>31<br>29<br>25<br>31<br>33<br>27<br>21<br>29<br>25<br>31<br>33<br>27<br>21<br>29<br>25<br>29<br>25<br>20<br>27<br>37<br>23<br>8<br>33<br>27<br>21<br>29<br>25<br>20<br>27<br>27<br>23<br>8<br>33<br>27<br>21<br>29<br>25<br>20<br>27<br>27<br>23<br>8<br>33<br>27<br>21<br>29<br>25<br>20<br>27<br>27<br>23<br>8<br>33<br>27<br>21<br>29<br>25<br>20<br>27<br>27<br>23<br>8<br>8<br>20<br>27<br>27<br>29<br>29<br>25<br>20<br>27<br>27<br>29<br>29<br>25<br>20<br>27<br>27<br>29<br>29<br>25<br>20<br>27<br>27<br>29<br>25<br>29<br>20<br>27<br>21<br>29<br>25<br>29<br>25<br>20<br>27<br>27<br>29<br>29<br>25<br>29<br>29<br>25<br>29<br>29<br>25<br>29<br>29<br>25<br>29<br>29<br>25<br>29<br>29<br>25<br>29<br>25<br>29<br>25<br>29<br>25<br>29<br>25<br>29<br>25<br>29<br>25<br>29<br>25<br>29<br>25<br>29<br>25<br>29<br>25<br>29<br>25<br>29<br>25<br>31<br>29<br>25<br>31<br>29<br>25<br>31<br>29<br>25<br>31<br>29<br>25<br>31<br>29<br>25<br>31<br>29<br>25<br>31<br>29<br>25<br>31<br>29<br>25<br>31<br>29<br>25<br>31<br>29<br>25<br>31<br>29<br>25<br>31<br>29<br>25<br>31<br>29<br>25<br>31<br>29<br>25<br>31<br>29<br>25<br>31<br>29<br>25<br>31<br>29<br>25<br>31<br>27<br>26<br>29<br>25<br>31<br>27<br>26<br>28<br>28<br>28<br>28<br>28<br>28<br>28<br>28<br>28<br>28<br>28<br>28<br>28   | Total           311           300           44           511           277           12           74           355           311           40           39           400           633           555           555           433           400           47           355           433           400           47           355           433           40           47           355           433           4335           355           355           433           355           355           355           355           355           355           36           375           38           39           305           355           36           375           375  | Weialit 2,5% 2,4% 3,5% 4,1% 2,2% 1,0% 2,2% 3,3% 3,1% 3,2% 3,3% 3,1% 3,1% 3,2% 3,3% 3,1% 3,5% 3,3% 3,2% 2,8% 2,0% 3,5% 3,3% 3,2% 2,8% 2,0% 3,5% 2,8% 2,0% 3,5% 2,8% 2,8% 2,0% 3,5% 2,8% 2,8% 2,8% 2,8% 2,8% 2,8% 2,8% 2,8   | Risk Difference<br>M-H, Fixed, 95% CI<br>0.16 [0.01, 0.31]<br>0.13 [-0.09, 0.35]<br>0.00 [-0.20, 0.20]<br>0.04 [-0.13, 0.21]<br>-0.07 [-0.27, 0.14]<br>0.25 [-0.06, 0.56]<br>0.28 [0.12, 0.43]<br>0.03 [-0.16, 0.22]<br>0.10 [-0.12, 0.32]<br>0.00 [-0.21, 0.21]<br>0.01 [-0.19, 0.21]<br>0.32 [0.05, 0.41]<br>0.32 [0.05, 0.41]<br>0.32 [0.05, 0.40]<br>0.05 [-0.09, 0.20]<br>0.19 [0.01, 0.37]<br>0.04 [-0.16, 0.24]<br>0.09 [-0.27, 0.24]<br>0.09 [-0.27, 0.24]<br>0.05 [-0.09, 0.20]<br>0.19 [0.01, 0.37]<br>0.04 [-0.16, 0.24]<br>0.09 [-0.27, 0.24]<br>0.09 [-0.22, 0.34]<br>0.21 [0.03, 0.39]<br>0.22 [0.05, 0.40]<br>0.20 [0.00, 0.43]<br>0.20 [0.00, 0.4   | 0.01 0.1 1 10 10 Favours [Kanglaite] Favours [Chemotherapy] Risk Difference M-H, Fixed, 95% Ci          |
| Test for overall effect<br>Study or Subgroup<br>Bao H 2019<br>Chen C 2018<br>Chen W 2018<br>Chen VZ 2003<br>Buan XG 2009<br>He LT 2017<br>LL 2016<br>Lang SG 2014<br>Lu F 2015<br>Long SG 2017<br>Mu G 2018<br>Sun SG 2012<br>Nang H 2016<br>Nang L 2016<br>Nang L 2016<br>Yang L 2018<br>Yang L 2018<br>Yang L 2018<br>Yang L 2018<br>Yang L 2018<br>Yang L 2018<br>Yang L 2018   | Z = 6.47<br>Kangla<br><u>Events</u><br>30<br>24<br>27<br>39<br>22<br>11<br>48<br>28<br>24<br>26<br>28<br>24<br>25<br>28<br>34<br>36<br>18<br>54<br>28<br>34<br>36<br>18<br>54<br>29<br>41<br>27<br>33<br>39<br>20<br>22<br>41<br>36<br>28<br>34<br>36<br>28<br>36<br>28<br>36<br>39<br>22<br>24<br>25<br>28<br>34<br>36<br>28<br>36<br>28<br>28<br>28<br>28<br>28<br>28<br>28<br>28<br>28<br>28   | - 30 (P < 0.0.<br>Total 31<br>30<br>31<br>30<br>44<br>45<br>12<br>28<br>41<br>28<br>35<br>51<br>55<br>55<br>55<br>55<br>55<br>55<br>55<br>55<br>5  | Chemothin Events  25 20 27 27 27 23 8 33 27 21 25 20 27 21 25 31 15 43 26 44 28 26 37 26 43 26 44 28 26 37 26 43 26 43 26 44 28 26 37 26 28 23 28 23 28 23 28 23 28 23 28 28 28 28 28 28 28 28 28 28 28 28 28   | Image: second | Weialitt<br>2.5%<br>2.4%<br>3.5%<br>4.1%<br>2.2%<br>3.3%<br>3.1%<br>3.2%<br>3.1%<br>3.1%<br>2.8%<br>4.4%<br>3.3%<br>3.8%<br>2.8%<br>3.3%<br>3.8%<br>2.8%<br>3.3%<br>3.8%<br>2.0%<br>3.5%<br>3.5%<br>3.5%<br>3.5%<br>3.5%<br>3.5%<br>3.5%<br>3.5  | Risk Difference<br>M-H, Fixed, 95% CI<br>0.16 [0.01, 0.31]<br>0.00 [-0.20, 0.20]<br>0.04 [-0.13, 0.21]<br>-0.07 [-0.27, 0.14]<br>0.25 [-0.06, 0.56]<br>0.28 [0.12, 0.43]<br>0.03 [-0.16, 0.22]<br>0.10 [-0.12, 0.32]<br>0.06 [-0.21, 0.21]<br>0.23 [0.05, 0.41]<br>0.23 [0.05, 0.41]<br>0.23 [0.05, 0.41]<br>0.23 [0.05, 0.41]<br>0.23 [0.05, 0.41]<br>0.33 [-0.22, 0.29]<br>0.17 [0.03, 0.32]<br>0.06 [-0.14, 0.25]<br>0.05 [-0.09, 0.20]<br>0.19 [0.01, 0.37]<br>0.04 [-0.16, 0.24]<br>0.03 [-0.22, 0.34]<br>0.03 [-0.22, 0.34]<br>0.05 [-0.09, 0.20]<br>0.45 [-0.17, 0.23]<br>0.06 [-0.22, 0.34]<br>0.21 [0.03, 0.39]<br>0.22 [-0.51, 0.18]<br>0.20 [0.00, 0.40]<br>0.15 [-0.06, 0.40]<br>0.15 [-0.06, 0.40]<br>0.45 [-0.22, 0.29]  | 0.01 0.1 1 10 10 Favours [Kanglaite] Favours [Chemotherapy] Risk Difference M.H. Fixed, 95% Cl          |
| Test for overall effect<br>Study or Subgroup<br>Bao H 2019<br>Chen V 2019<br>Chen V 2018<br>Chen V 2018<br>Chen V 2018<br>Chen V 2017<br>Huang ZB 2010<br>HV 2017<br>Huang ZB 2010<br>HV 2017<br>JL 2012<br>JND 2018<br>Lang JG 2014<br>Ju F 2015<br>Long SG 2017<br>Ju Z 2018<br>Sun SQ 2012<br>Nang LV 2018<br>Sun SQ 2012<br>Nang V 2017<br>(an QH 2018<br>Sun SQ 2014<br>Nang V 2017<br>(an QH 2018<br>Sun SQ 2014<br>Nang V 2017<br>(an QH 2018<br>(ang L 2018<br>(ang L 2018<br>(ang L 2018<br>(ang L 2018<br>(ang SJ 2003<br>(ao J 2017<br>(an QH 2018<br>(ang SJ 2003<br>(ao J 2017<br>(ang L 2018   | Z = 6.47<br>Kangla<br>Events<br>30<br>24<br>27<br>39<br>22<br>11<br>48<br>28<br>24<br>26<br>28<br>34<br>366<br>18<br>54<br>29<br>41<br>18<br>54<br>29<br>41<br>13<br>37<br>33<br>39<br>39<br>29<br>41<br>27<br>29<br>41<br>27<br>29<br>41<br>36<br>29<br>47<br>27<br>29<br>29<br>47<br>27<br>29<br>29<br>29<br>47<br>27<br>29<br>29<br>29<br>29<br>29<br>29<br>29<br>29<br>29<br>29   | Solution           Solution <t< td=""><td>Chemothin<br/>Events<br/>25<br/>20<br/>27<br/>37<br/>23<br/>8<br/>8<br/>33<br/>27<br/>21<br/>25<br/>28<br/>25<br/>31<br/>15<br/>43<br/>26<br/>44<br/>428<br/>26<br/>44<br/>28<br/>26<br/>42<br/>27<br/>21<br/>15<br/>43<br/>26<br/>25<br/>31<br/>15<br/>43<br/>26<br/>25<br/>29<br/>25<br/>29<br/>25<br/>29<br/>25<br/>20<br/>27<br/>23<br/>23<br/>26<br/>20<br/>27<br/>27<br/>23<br/>23<br/>26<br/>20<br/>27<br/>27<br/>23<br/>23<br/>26<br/>20<br/>27<br/>23<br/>26<br/>20<br/>27<br/>27<br/>23<br/>20<br/>27<br/>23<br/>20<br/>27<br/>23<br/>20<br/>27<br/>23<br/>20<br/>27<br/>23<br/>20<br/>27<br/>23<br/>20<br/>27<br/>23<br/>20<br/>27<br/>23<br/>20<br/>27<br/>23<br/>20<br/>27<br/>23<br/>20<br/>27<br/>23<br/>20<br/>27<br/>23<br/>26<br/>20<br/>27<br/>25<br/>20<br/>20<br/>27<br/>23<br/>20<br/>25<br/>20<br/>20<br/>27<br/>25<br/>20<br/>20<br/>27<br/>25<br/>20<br/>20<br/>27<br/>25<br/>20<br/>20<br/>27<br/>25<br/>20<br/>20<br/>27<br/>25<br/>20<br/>20<br/>20<br/>27<br/>25<br/>20<br/>20<br/>25<br/>20<br/>20<br/>25<br/>20<br/>20<br/>25<br/>20<br/>20<br/>25<br/>20<br/>20<br/>25<br/>20<br/>20<br/>25<br/>20<br/>20<br/>25<br/>20<br/>25<br/>20<br/>20<br/>25<br/>20<br/>20<br/>25<br/>20<br/>20<br/>25<br/>20<br/>25<br/>20<br/>25<br/>20<br/>25<br/>20<br/>25<br/>20<br/>25<br/>20<br/>25<br/>20<br/>25<br/>20<br/>25<br/>25<br/>20<br/>25<br/>20<br/>25<br/>20<br/>25<br/>20<br/>25<br/>25<br/>20<br/>25<br/>25<br/>20<br/>25<br/>25<br/>20<br/>25<br/>25<br/>20<br/>25<br/>25<br/>20<br/>25<br/>25<br/>20<br/>25<br/>25<br/>20<br/>25<br/>25<br/>20<br/>25<br/>25<br/>20<br/>26<br/>25<br/>20<br/>26<br/>25<br/>20<br/>27<br/>23<br/>27<br/>25<br/>20<br/>26<br/>25<br/>20<br/>20<br/>27<br/>20<br/>20<br/>20<br/>20<br/>20<br/>20<br/>20<br/>20<br/>20<br/>20<br/>20<br/>20<br/>20</td><td>Prapy           Total           31           300           44           51           27           12           54           35           31           40           39           40           39           40           30           355           43           40           63           355           43           40           63           355           43           40           63           355           43           40           63           355           43           40           55           43           40           35           36           49           35           67           67</td><td>Weight 2,5% 2,4% 3,5% 4,1% 2,2% 4,3% 2,2% 4,3% 2,8% 3,1% 3,1% 3,2% 3,1% 3,1% 3,2% 3,3% 3,2% 2,8% 2,0% 3,5% 3,3% 3,2% 2,8% 2,0% 3,9% 2,8% 2,0% 3,9% 2,8% 2,0% 3,9% 2,8% 2,0% 3,9% 2,2% 4,0% 3,0% 2,0% 3,0\% 2,0\% 2,0\% 2,0\% 2,0\% 2,0\% 2,0\% 2,0\% 2</td><td>Risk Difference<br/>M-H, Fixed, 95% CI<br/>0.18 [0.01, 0.35]<br/>0.00 [-0.20, 0.20]<br/>0.04 [-0.13, 0.21]<br/>-0.07 [-0.27, 0.14]<br/>0.25 [-0.06, 0.56]<br/>0.28 [0.12, 0.43]<br/>0.03 [-0.16, 0.22]<br/>0.10 [-0.21, 0.32]<br/>0.00 [-0.21, 0.21]<br/>0.01 [-0.19, 0.21]<br/>0.33 [-0.03, 0.28]<br/>0.03 [-0.22, 0.29]<br/>0.17 [0.03, 0.32]<br/>0.05 [-0.09, 0.20]<br/>0.19 [0.01, 0.37]<br/>0.04 [-0.16, 0.24]<br/>0.03 [-0.27, 0.24]<br/>0.03 [-0.17, 0.23]<br/>0.06 [-0.14, 0.25]<br/>0.05 [-0.09, 0.20]<br/>0.19 [0.01, 0.37]<br/>0.4 [-0.16, 0.24]<br/>0.03 [-0.17, 0.23]<br/>0.06 [-0.22, 0.34]<br/>0.22 [0.05, 0.40]<br/>0.22 [0.05, 0.40]<br/>0.20 [-0.06, 0.36]<br/>0.4 [-0.12, 0.26]<br/>0.4 [-0.12, 0</td><td>0.01 0.1 1 10 10 Favours [Kanglaite] Favours [Chemotherapy] Risk Difference M.H. Fixed, 95% Ci</td></t<> | Chemothin<br>Events<br>25<br>20<br>27<br>37<br>23<br>8<br>8<br>33<br>27<br>21<br>25<br>28<br>25<br>31<br>15<br>43<br>26<br>44<br>428<br>26<br>44<br>28<br>26<br>42<br>27<br>21<br>15<br>43<br>26<br>25<br>31<br>15<br>43<br>26<br>25<br>29<br>25<br>29<br>25<br>29<br>25<br>20<br>27<br>23<br>23<br>26<br>20<br>27<br>27<br>23<br>23<br>26<br>20<br>27<br>27<br>23<br>23<br>26<br>20<br>27<br>23<br>26<br>20<br>27<br>27<br>23<br>20<br>27<br>23<br>20<br>27<br>23<br>20<br>27<br>23<br>20<br>27<br>23<br>20<br>27<br>23<br>20<br>27<br>23<br>20<br>27<br>23<br>20<br>27<br>23<br>20<br>27<br>23<br>20<br>27<br>23<br>20<br>27<br>23<br>26<br>20<br>27<br>25<br>20<br>20<br>27<br>23<br>20<br>25<br>20<br>20<br>27<br>25<br>20<br>20<br>27<br>25<br>20<br>20<br>27<br>25<br>20<br>20<br>27<br>25<br>20<br>20<br>27<br>25<br>20<br>20<br>20<br>27<br>25<br>20<br>20<br>25<br>20<br>20<br>25<br>20<br>20<br>25<br>20<br>20<br>25<br>20<br>20<br>25<br>20<br>20<br>25<br>20<br>20<br>25<br>20<br>25<br>20<br>20<br>25<br>20<br>20<br>25<br>20<br>20<br>25<br>20<br>25<br>20<br>25<br>20<br>25<br>20<br>25<br>20<br>25<br>20<br>25<br>20<br>25<br>20<br>25<br>25<br>20<br>25<br>20<br>25<br>20<br>25<br>20<br>25<br>25<br>20<br>25<br>25<br>20<br>25<br>25<br>20<br>25<br>25<br>20<br>25<br>25<br>20<br>25<br>25<br>20<br>25<br>25<br>20<br>25<br>25<br>20<br>25<br>25<br>20<br>26<br>25<br>20<br>26<br>25<br>20<br>27<br>23<br>27<br>25<br>20<br>26<br>25<br>20<br>20<br>27<br>20<br>20<br>20<br>20<br>20<br>20<br>20<br>20<br>20<br>20<br>20<br>20<br>20  | Prapy           Total           31           300           44           51           27           12           54           35           31           40           39           40           39           40           30           355           43           40           63           355           43           40           63           355           43           40           63           355           43           40           63           355           43           40           55           43           40           35           36           49           35           67           67   | Weight 2,5% 2,4% 3,5% 4,1% 2,2% 4,3% 2,2% 4,3% 2,8% 3,1% 3,1% 3,2% 3,1% 3,1% 3,2% 3,3% 3,2% 2,8% 2,0% 3,5% 3,3% 3,2% 2,8% 2,0% 3,9% 2,8% 2,0% 3,9% 2,8% 2,0% 3,9% 2,8% 2,0% 3,9% 2,2% 4,0% 3,0% 2,0% 3,0\% 2,0\% 2,0\% 2,0\% 2,0\% 2,0\% 2,0\% 2,0\% 2   | Risk Difference<br>M-H, Fixed, 95% CI<br>0.18 [0.01, 0.35]<br>0.00 [-0.20, 0.20]<br>0.04 [-0.13, 0.21]<br>-0.07 [-0.27, 0.14]<br>0.25 [-0.06, 0.56]<br>0.28 [0.12, 0.43]<br>0.03 [-0.16, 0.22]<br>0.10 [-0.21, 0.32]<br>0.00 [-0.21, 0.21]<br>0.01 [-0.19, 0.21]<br>0.33 [-0.03, 0.28]<br>0.03 [-0.22, 0.29]<br>0.17 [0.03, 0.32]<br>0.05 [-0.09, 0.20]<br>0.19 [0.01, 0.37]<br>0.04 [-0.16, 0.24]<br>0.03 [-0.27, 0.24]<br>0.03 [-0.17, 0.23]<br>0.06 [-0.14, 0.25]<br>0.05 [-0.09, 0.20]<br>0.19 [0.01, 0.37]<br>0.4 [-0.16, 0.24]<br>0.03 [-0.17, 0.23]<br>0.06 [-0.22, 0.34]<br>0.22 [0.05, 0.40]<br>0.22 [0.05, 0.40]<br>0.20 [-0.06, 0.36]<br>0.4 [-0.12, 0.26]<br>0.4 [-0.12, 0   | 0.01 0.1 1 10 10 Favours [Kanglaite] Favours [Chemotherapy] Risk Difference M.H. Fixed, 95% Ci          |
| Herefogenerity, Chr =           Fest for overall effect           Study or Suboroug           Bao H 2019           Chen V2018           Chen V2009           de LT 2017           Huang ZB 2010           ia JN 2018           Lang J 2018           Lang SG 2014           Ju F 2019           Ju JA 2016           Ju Y 2015           Jong SG 2017           Alua 2018           Sun SQ 2012           Nang H 2016           Nang Y 2017           (an QH 2018           (an QH 2018           (ang SJ 2003           (ao J 2017           (ang SJ 2003           (ao J 2017           (ao J 2017           (ang SJ 2003           (ao J 2017           (ao J 2017           (ao J 2017   | Z = 6.47<br>Kangla<br><u>Events</u><br>30<br>24<br>30<br>24<br>27<br>39<br>22<br>24<br>36<br>18<br>28<br>24<br>26<br>36<br>18<br>54<br>26<br>36<br>18<br>54<br>29<br>29<br>41<br>36<br>29<br>47<br>36<br>29<br>47<br>36<br>29<br>47<br>36<br>29<br>36<br>29<br>36<br>29<br>36<br>29<br>36<br>29<br>36<br>29<br>36<br>29<br>36<br>29<br>36<br>29<br>36<br>29<br>36<br>29<br>36<br>29<br>36<br>36<br>29<br>36<br>36<br>36<br>36<br>36<br>36<br>36<br>36<br>36<br>36   | - 30 (P < 0.0.<br>Total 31<br>30 0 44<br>31 30 0 44<br>35 31<br>28 8<br>12 2<br>35 31<br>31 30<br>39 40<br>33 35<br>55 55<br>43 35<br>55 54<br>34 42<br>47 35<br>24 43<br>36 35<br>55 54<br>35 55<br>24 43<br>36 35<br>27 24<br>47 35<br>24 43<br>36 35<br>27 24<br>47 35<br>27 27<br>47 35<br>27 24<br>47 35<br>27 27<br>47 35<br>27 27<br>27 35<br>27 27<br>27 35<br>27 27<br>27 35<br>27 35  | Chemothin<br>Events<br>25<br>20<br>27<br>37<br>23<br>8<br>33<br>27<br>23<br>8<br>33<br>27<br>21<br>25<br>29<br>25<br>31<br>15<br>43<br>26<br>44<br>428<br>26<br>37<br>21<br>15<br>43<br>26<br>44<br>428<br>26<br>37<br>21<br>29<br>25<br>31<br>29<br>25<br>31<br>29<br>25<br>31<br>29<br>25<br>31<br>29<br>25<br>31<br>29<br>25<br>20<br>27<br>37<br>23<br>8<br>8<br>33<br>27<br>21<br>29<br>25<br>20<br>27<br>27<br>23<br>8<br>8<br>33<br>27<br>21<br>29<br>25<br>20<br>27<br>27<br>23<br>8<br>8<br>33<br>27<br>21<br>29<br>25<br>20<br>27<br>27<br>23<br>8<br>8<br>33<br>27<br>29<br>25<br>20<br>27<br>27<br>29<br>29<br>25<br>20<br>27<br>27<br>27<br>29<br>25<br>20<br>27<br>27<br>29<br>25<br>29<br>25<br>20<br>27<br>27<br>29<br>25<br>29<br>20<br>27<br>27<br>29<br>25<br>29<br>25<br>29<br>25<br>29<br>25<br>29<br>25<br>29<br>25<br>29<br>25<br>29<br>25<br>29<br>25<br>29<br>25<br>29<br>25<br>29<br>25<br>29<br>25<br>29<br>25<br>29<br>25<br>29<br>25<br>29<br>25<br>29<br>25<br>29<br>25<br>29<br>25<br>29<br>25<br>29<br>25<br>29<br>25<br>29<br>25<br>29<br>25<br>31<br>27<br>29<br>25<br>31<br>27<br>29<br>25<br>29<br>25<br>31<br>25<br>29<br>25<br>31<br>27<br>29<br>25<br>31<br>27<br>29<br>25<br>31<br>27<br>29<br>25<br>31<br>29<br>25<br>31<br>27<br>29<br>25<br>31<br>29<br>25<br>31<br>29<br>25<br>31<br>27<br>29<br>25<br>31<br>29<br>25<br>31<br>27<br>29<br>25<br>31<br>27<br>29<br>25<br>31<br>27<br>26<br>28<br>28<br>26<br>26<br>28<br>28<br>28<br>27<br>29<br>25<br>31<br>27<br>26<br>28<br>28<br>28<br>28<br>28<br>26<br>26<br>29<br>25<br>29<br>25<br>29<br>25<br>29<br>25<br>29<br>25<br>29<br>25<br>29<br>25<br>29<br>25<br>29<br>25<br>29<br>25<br>29<br>25<br>29<br>25<br>29<br>25<br>29<br>25<br>29<br>25<br>29<br>25<br>29<br>25<br>29<br>26<br>26<br>27<br>29<br>26<br>27<br>29<br>26<br>26<br>27<br>27<br>29<br>25<br>29<br>29<br>25<br>29<br>25<br>29<br>25<br>29<br>20<br>25<br>29<br>25<br>29<br>25<br>29<br>25<br>29<br>25<br>29<br>25<br>29<br>25<br>29<br>25<br>29<br>25<br>29<br>25<br>29<br>25<br>29<br>25<br>29<br>25<br>29<br>25<br>29<br>25<br>29<br>25<br>29<br>29<br>25<br>29<br>25<br>29<br>25<br>29<br>25<br>29<br>25<br>25<br>25<br>29<br>25<br>25<br>25<br>25<br>25<br>25<br>25<br>25<br>25<br>25<br>25<br>25<br>25 | Total           311           300           44           511           277           12           74           355           331           40           399           400           633           555           433           555           433           369           255           433           369           255           265           433           369           255           261           433           355           266           667           400           410           420           355           266           677           400  | Weialit 2,5% 2,4% 3,5% 4,1% 2,2% 1,0% 2,2% 3,3% 3,1% 3,2% 3,3% 3,1% 3,1% 3,1% 3,1% 3,2% 4,4% 3,5% 3,3% 3,3% 3,3% 3,3% 3,3% 3,3% 3,5% 2,8% 2,0% 3,5% 2,5% 3,2% 2,5% 3,2% 2,5% 3,2% 2,5% 3,2% 2,5% 3,2% 2,5% 3,2% 2,5% 3,2% 2,5% 3,2% 2,5% 3,2% 2,5% 3,2% 2,5% 3,2% 2,5% 3,2% 2,5% 3,2% 2,5% 3,2% 2,5% 3,2% 2,5% 3,2% 2,5% 3,2\% 2,5\% 2,5\% 3,2\% 2,5\% 2,5\% 2,5\% 2,5\% 2,5\% 2,5\% 2,5\% 2 | Risk Difference<br>M-H, Fixed, 95% CI<br>0.16 [0.01, 0.31]<br>0.13 [-0.09, 0.35]<br>0.00 [-0.20, 0.20]<br>0.04 [-0.13, 0.21]<br>-0.07 [-0.27, 0.14]<br>0.25 [-0.06, 0.56]<br>0.28 [0.12, 0.43]<br>0.03 [-0.16, 0.22]<br>0.10 [-0.12, 0.32]<br>0.00 [-0.21, 0.21]<br>0.01 [-0.19, 0.21]<br>0.03 [-0.20, 0.24]<br>0.03 [-0.20, 0.24]<br>0.03 [-0.20, 0.24]<br>0.03 [-0.22, 0.24]<br>0.04 [-0.16, 0.24]<br>0.05 [-0.09, 0.20]<br>0.19 [0.01, 0.37]<br>0.04 [-0.17, 0.23]<br>0.06 [-0.22, 0.34]<br>0.21 [0.03, 0.39]<br>0.22 [0.05, 0.40]<br>0.22 [-0.14, 0.18]<br>0.20 [-0.00, 0.40]<br>0.15 [-0.06, 0.36]<br>0.4 [-0.12, 0.20]<br>0.03 [-0.11, 0.16]<br>0.24 [-0.12, 0.20]<br>0.03 [-0.11, 0.16]<br>0.26 [-0.20, 0.40]<br>0.25 [-0.00, 0           | 0.01 0.1 1 10 10 Favours [Kanglaite] Favours [Chemotherapy]  Risk Difference M.H. Fixed, 95% Ci         |
| Test for overall effect<br>Test for overall effect<br>Study or Subgroup<br>Bao H 2019<br>Chen C 2018<br>Chen W 2016<br>Chen Y 2018<br>Chen YZ 2003<br>Suan X0 2009<br>He LT 2017<br>Huang ZB 2010<br>Jia JN 2018<br>Li HY 2017<br>Li L 2017<br>Huang ZB 2010<br>Jia JN 2018<br>Liang SC 2014<br>Liu F 2019<br>Liu SC 2014<br>Liu Y 2015<br>Long SG 2017<br>Mu G 2018<br>Sun SQ 2012<br>Nang HY 2016<br>Nang L 2014<br>Nang Y 2017<br>(an GH 2018<br>(ang L 2017<br>(ang L 2018<br>(ang L 2018)<br>(ang L             | Z = 6.47<br>Kangla<br><u>Events</u><br>30<br>24<br>27<br>39<br>22<br>11<br>48<br>28<br>24<br>26<br>28<br>24<br>26<br>28<br>34<br>36<br>29<br>41<br>27<br>36<br>39<br>30<br>22<br>47<br>36<br>37<br>39<br>30<br>22<br>28<br>34<br>36<br>28<br>28<br>28<br>28<br>28<br>28<br>28<br>28<br>28<br>28   | - 50 (P < 0.0.<br>(P < 0.0.<br>31 (P < 0.0.<br>31 30 (P < 0.0.<br>32 (P < 0.0.<br>31 30 (P < 0.0.<br>31   | Chemothin Events  25 20 27 27 23 8 33 27 21 25 28 25 31 15 15 43 26 37 26 37 26 37 26 37 26 37 26 37 26 37 26 37 26 37 26 37 26 37 26 37 26 37 26 37 26 37 36 44 28 26 37 26 37 26 37 36 42 38 38 39 42 35 38 39 42 35 38 38 38 38 38 38 38 38 38 38 38 38 38   | Total           311           300           44           511           277           12           277           12           355           311           41           40           39           400           303           555           555           555           555           43           36           355           265           433           36           355           265           67           60           60           60           61           62           67           60           60           60           60  | Weialitt<br>2.5%<br>2.4%<br>3.5%<br>4.1%<br>2.2%<br>1.0%<br>4.3%<br>2.8%<br>3.1%<br>3.1%<br>3.1%<br>3.1%<br>3.1%<br>3.1%<br>3.1%<br>3.3%<br>3.3  | Risk Difference<br>M-H, Fixed, 95% CI<br>0.16 [0.01, 0.31]<br>0.13 [-0.09, 0.35]<br>0.00 [-0.20, 0.20]<br>0.04 [-0.13, 0.21]<br>-0.07 [-0.27, 0.14]<br>0.25 [-0.06, 0.56]<br>0.28 [0.12, 0.43]<br>0.03 [-0.16, 0.22]<br>0.10 [-0.21, 0.21]<br>0.23 [0.05, 0.41]<br>0.13 [-0.03, 0.28]<br>0.03 [-0.29, 0.29]<br>0.17 [0.03, 0.32]<br>0.06 [-0.22, 0.29]<br>0.17 [0.03, 0.32]<br>0.05 [-0.09, 0.20]<br>0.19 [-0.01, 0.37]<br>0.04 [-0.16, 0.24]<br>0.03 [-0.22, 0.24]<br>0.05 [-0.09, 0.20]<br>0.05 [-0.22, 0.34]<br>0.21 [0.03, 0.39]<br>0.22 [-0.05, 0.40]<br>0.22 [-0.05, 0.40]<br>0.23 [-0.14, 0.18]<br>0.20 [0.00, 0.40]<br>0.34 [-0.12, 0.20]<br>0.03 [-0.11, 0.16]<br>0.23 [0.08, 0.38]<br>0.30 [-0.11, 0.16]<br>0.32 [0.08, 0.38]<br>0.30 [-0.11, 0.16]<br>0.33 [-0.11, 0.16]<br>0.34 [-0.11, 0.20]<br>0.34 [-0.11, 0.20]<br>0.         | 0.01 0.1 1 10 10 Favours [Kanglaite] Favours [Chemotherapy] Risk Difference M.H. Fixed, 95% Cl          |
| Test for overall effect<br>Test for overall effect<br>Study or Subgroup<br>Bao H 2019<br>Chen C 2018<br>Chen W 2016<br>Chen YZ 2003<br>Guan XG 2009<br>He LT 2017<br>Huang ZB 2010<br>Jia JN 2018<br>Li HY 2017<br>Li L 2012<br>Li XD 2016<br>Liang S 2014<br>Liu F 2019<br>Liu JG 2011<br>Liu Y 2015<br>Long SG 2017<br>Wu G 2018<br>Sun SQ 2012<br>Nang HY 2016<br>Nang L 2018<br>Nang L 2018<br>Yang Y 2017<br>Yang L 2018<br>Yang L 2018<br>Yang L 2018<br>Yang L 2018<br>Yang Y 2017<br>Yang Y 2019<br>Yang Y 2017<br>Yang Yang Y 2017<br>Yang Yang Yang Yang Yang    | Z = 6.47<br>Kangla<br><u>Events</u><br>30<br>24<br>27<br>39<br>22<br>11<br>48<br>28<br>24<br>26<br>28<br>24<br>26<br>28<br>34<br>36<br>18<br>54<br>28<br>36<br>18<br>54<br>29<br>41<br>27<br>36<br>29<br>41<br>37<br>36<br>29<br>41<br>36<br>29<br>41<br>36<br>28<br>36<br>28<br>36<br>28<br>36<br>28<br>36<br>28<br>36<br>28<br>36<br>28<br>36<br>28<br>36<br>28<br>36<br>28<br>28<br>28<br>28<br>28<br>28<br>28<br>28<br>28<br>28   | - 30 (P < 0.0<br>(P < 0.0<br>31<br>30<br>30<br>44<br>451<br>28<br>31<br>28<br>32<br>31<br>32<br>34<br>35<br>55<br>55<br>55<br>55<br>55<br>55<br>55<br>55<br>55   | Chemothin Events  25 20 27 27 27 23 8 33 27 21 25 29 25 31 15 15 43 26 44 28 26 37 26 43 26 37 26 12 28 25 31 19 42 28 25 33 19 42 25 38 20   | Prapy           Total           31           300           44           51           27           12           355           311           40           39           400           300           633           555           555           555           555           433           40           477           355           555           555           555           565           555           433           40           477           355           255           433           36           667           40           600           500   | Weialitt<br>2.5%<br>2.4%<br>3.5%<br>2.2%<br>1.0%<br>4.3%<br>2.8%<br>3.1%<br>3.1%<br>3.1%<br>3.1%<br>3.1%<br>3.1%<br>3.1%<br>3.1  | Risk Difference<br>M-H, Fixed, 95% CI<br>0.16 [0.01, 0.31]<br>0.13 [-0.09, 0.35]<br>0.00 [-0.20, 0.20]<br>0.04 [-0.13, 0.21]<br>-0.07 [-0.27, 0.14]<br>0.25 [-0.06, 0.56]<br>0.28 [0.12, 0.43]<br>0.03 [-0.16, 0.22]<br>0.10 [-0.21, 0.21]<br>0.23 [0.05, 0.41]<br>0.23 [0.05, 0.41]<br>0.23 [0.05, 0.41]<br>0.32 [0.05, 0.41]<br>0.32 [0.05, 0.41]<br>0.34 [-0.16, 0.22]<br>0.05 [-0.09, 0.20]<br>0.17 [0.03, 0.32]<br>0.06 [-0.22, 0.29]<br>0.17 [0.03, 0.32]<br>0.05 [-0.09, 0.20]<br>0.19 [0.01, 0.37]<br>0.04 [-0.16, 0.24]<br>0.03 [-0.17, 0.23]<br>0.06 [-0.22, 0.34]<br>0.22 [0.05, 0.40]<br>0.22 [0.05, 0.40]<br>0.22 [0.05, 0.40]<br>0.24 [-0.14, 0.18]<br>0.20 [0.00, 0.40]<br>0.34 [-0.14, 0.12]<br>0.24 [-0.14, 0.18]<br>0.20 [0.00, 0.40]<br>0.34 [-0.11, 0.16]<br>0.23 [0.08, 0.38]<br>0.30 [0.11, 0.49]  | 0.01 0.1 1 10 10 Favours [Kanglaite] Favours [Chemotherapy] Risk Difference M.H. Fixed, 95% Cl          |
| Test for overall effect<br>Test for overall effect<br>Study or Subgroup<br>Bao H 2019<br>Chen C 2018<br>Chen YZ 2003<br>Ghen YZ 2017<br>Hang ZB 2010<br>Ja JN 2018<br>Li HY 2017<br>Li L 2012<br>Li XD 2016<br>Liang SC 2014<br>Jiu Y 2015<br>Long SG 2017<br>Wu G 2018<br>Sum SQ 2012<br>Nang HY 2016<br>Nang L 2018<br>Yang Y 2017<br>Yang L 2018<br>Yang L 2018<br>Yang L 2018<br>Yang L 2018<br>Yang Y 2017<br>Yang Y 2 | Z = 6.47<br>Kangla<br><u>Events</u><br>30<br>24<br>27<br>39<br>22<br>11<br>48<br>28<br>24<br>25<br>28<br>34<br>36<br>18<br>54<br>28<br>34<br>36<br>18<br>54<br>28<br>34<br>36<br>29<br>41<br>27<br>36<br>29<br>41<br>37<br>36<br>29<br>41<br>36<br>26<br>28<br>34<br>36<br>28<br>36<br>28<br>36<br>28<br>36<br>28<br>28<br>28<br>28<br>28<br>28<br>28<br>28<br>28<br>28   | - 30 (P < 0.0.<br>P < 0.0.<br>31 300<br>34 4<br>51 28<br>32 54<br>54 51<br>28 8<br>12 2<br>54 51<br>31 30<br>41 41<br>35 55<br>55 55<br>55 55<br>55 55<br>55 55<br>55 55<br>55 55<br>55 55<br>55 70<br>0 43<br>42 43<br>36 43<br>36 43<br>36 43<br>37 45<br>24 43<br>36 60<br>50 50<br>1249  | Chemothu<br>Events<br>25<br>20<br>27<br>27<br>23<br>8<br>33<br>27<br>21<br>25<br>20<br>27<br>23<br>8<br>33<br>27<br>21<br>25<br>20<br>27<br>21<br>25<br>20<br>27<br>21<br>25<br>20<br>27<br>21<br>25<br>20<br>27<br>21<br>25<br>20<br>27<br>21<br>25<br>20<br>27<br>21<br>25<br>20<br>27<br>21<br>25<br>20<br>27<br>21<br>25<br>20<br>27<br>21<br>25<br>20<br>27<br>21<br>25<br>20<br>27<br>21<br>25<br>20<br>27<br>21<br>25<br>20<br>27<br>21<br>25<br>20<br>27<br>21<br>25<br>20<br>27<br>21<br>25<br>26<br>26<br>27<br>21<br>25<br>26<br>26<br>27<br>21<br>25<br>26<br>26<br>27<br>21<br>25<br>26<br>26<br>26<br>27<br>21<br>25<br>26<br>26<br>26<br>27<br>21<br>25<br>26<br>25<br>31<br>15<br>43<br>26<br>43<br>26<br>43<br>26<br>43<br>26<br>43<br>26<br>43<br>26<br>43<br>26<br>43<br>26<br>43<br>26<br>43<br>26<br>43<br>26<br>43<br>26<br>43<br>26<br>43<br>26<br>25<br>31<br>15<br>43<br>26<br>25<br>31<br>26<br>25<br>31<br>26<br>25<br>31<br>26<br>25<br>31<br>26<br>25<br>31<br>26<br>25<br>31<br>26<br>25<br>31<br>26<br>25<br>31<br>26<br>25<br>31<br>26<br>25<br>31<br>26<br>25<br>31<br>26<br>25<br>31<br>26<br>25<br>31<br>26<br>25<br>31<br>26<br>25<br>31<br>26<br>25<br>31<br>26<br>25<br>31<br>26<br>25<br>31<br>26<br>25<br>31<br>26<br>25<br>31<br>26<br>25<br>31<br>26<br>25<br>31<br>26<br>25<br>31<br>26<br>25<br>31<br>26<br>25<br>31<br>26<br>25<br>31<br>20<br>25<br>25<br>25<br>31<br>26<br>25<br>25<br>25<br>25<br>25<br>25<br>25<br>25<br>25<br>25  | Prapy           Total           31           300           44           51           27           12           355           311           40           39           400           39           400           200           633           555           555           555           433           40           47           355           255           555           565           555           433           40           47           36           47           36           57           433           49           355           266           67           40           60           50           50           50           50           50           50           50   | Weialitt<br>2.5%<br>2.4%<br>3.5%<br>4.1%<br>2.2%<br>1.0%<br>4.3%<br>3.1%<br>3.1%<br>3.1%<br>3.1%<br>3.1%<br>3.1%<br>3.1%<br>3  | Risk Difference<br>M-H, Fixed, 95% CI<br>0.16 [0.01, 0.31]<br>0.13 [-0.09, 0.35]<br>0.00 [-0.20, 0.20]<br>0.04 [-0.13, 0.21]<br>-0.07 [-0.27, 0.14]<br>0.25 [-0.06, 0.56]<br>0.28 [0.12, 0.43]<br>0.03 [-0.16, 0.22]<br>0.10 [-0.21, 0.21]<br>0.23 [0.05, 0.41]<br>0.23 [0.05, 0.41]<br>0.23 [0.05, 0.41]<br>0.32 [0.05, 0.41]<br>0.32 [0.05, 0.41]<br>0.34 [-0.16, 0.22]<br>0.05 [-0.09, 0.20]<br>0.05 [-0.09, 0.20]<br>0.05 [-0.09, 0.20]<br>0.05 [-0.09, 0.20]<br>0.05 [-0.22, 0.34]<br>0.06 [-0.22, 0.34]<br>0.06 [-0.22, 0.34]<br>0.22 [0.05, 0.40]<br>0.22 [0.05, 0.40]<br>0.22 [0.05, 0.40]<br>0.22 [0.05, 0.40]<br>0.24 [-0.14, 0.18]<br>0.20 [-0.00, 0.40]<br>0.15 [-0.06, 0.38]<br>0.36 [0.11, 0.16]<br>0.23 [0.08, 0.38]<br>0.30 [0.11, 0.49]<br>0.11 [0.08, 0.15]  | 0.01 0.1 1 10 10<br>Favours [Kanglaite] Favours [Chemotherapy]<br>Risk Difference<br>M-H, Fixed, 95% Cl |
| Test for overall effect<br>Test for overall effect<br>Study or Subgroup<br>Bao H 2019<br>Chen C 2018<br>Chen V 2016<br>Chen V 2017<br>Huang ZB 2010<br>Jia JN 2018<br>LiAng ZB 2010<br>Jia JN 2018<br>LiAng SG 2014<br>Li V 2016<br>Liang J 2018<br>Liang SG 2014<br>Liu F 2019<br>Liu XH 2016<br>Liu XH 2016<br>Liu XH 2016<br>Ju Y 2017<br>Sun SQ 2012<br>Nang L 2014<br>Nang V 2017<br>Yang MY 2016<br>Sun SQ 2012<br>Nang L 2014<br>Nang Y 2017<br>Yang L 2018<br>Sun SQ 2012<br>Nang L 2018<br>Sun SQ 2012<br>Nang L 2018<br>Sun SQ 2012<br>Nang L 2014<br>Nang Y 2017<br>Yang J 2018<br>Sun SQ 2012<br>Nang L 2014<br>Nang Y 2017<br>Yang J 2018<br>Yang L 2018<br>Coll 2015<br>Zhang MM 2019<br>Total (95% CI)<br>Total events  | Z = 6.47<br>Kangla<br>Events<br>30<br>24<br>27<br>39<br>22<br>11<br>14<br>48<br>28<br>24<br>26<br>28<br>34<br>366<br>18<br>54<br>47<br>36<br>29<br>41<br>27<br>13<br>37<br>33<br>39<br>30<br>22<br>47<br>36<br>29<br>41<br>27<br>29<br>41<br>27<br>29<br>41<br>27<br>29<br>41<br>29<br>41<br>27<br>29<br>41<br>29<br>41<br>29<br>41<br>29<br>41<br>29<br>41<br>29<br>41<br>29<br>41<br>29<br>41<br>29<br>41<br>29<br>29<br>41<br>29<br>41<br>29<br>41<br>29<br>41<br>29<br>41<br>29<br>41<br>29<br>41<br>29<br>41<br>29<br>41<br>29<br>41<br>29<br>41<br>29<br>41<br>29<br>41<br>29<br>41<br>29<br>41<br>29<br>41<br>29<br>41<br>29<br>41<br>29<br>41<br>29<br>41<br>27<br>30<br>30<br>29<br>41<br>27<br>29<br>41<br>27<br>30<br>30<br>29<br>41<br>27<br>30<br>30<br>29<br>41<br>47<br>30<br>30<br>29<br>41<br>41<br>27<br>31<br>33<br>39<br>30<br>20<br>20<br>21<br>41<br>25<br>26<br>28<br>28<br>28<br>29<br>41<br>47<br>31<br>33<br>39<br>30<br>20<br>20<br>21<br>41<br>27<br>31<br>33<br>39<br>30<br>20<br>20<br>41<br>47<br>31<br>37<br>33<br>39<br>30<br>20<br>20<br>41<br>47<br>31<br>37<br>30<br>30<br>20<br>20<br>41<br>47<br>31<br>37<br>30<br>30<br>20<br>20<br>47<br>35<br>35<br>55<br>47<br>47<br>35<br>35<br>35<br>55<br>47<br>35<br>35<br>35<br>35<br>35<br>35<br>35<br>35<br>35<br>35 | Solution           Total           31           30           44           51           28           12           28           12           24           35           31           38           390           40           355           43           355           43           355           43           42           47           35           244           43           42           36           255           700           400           50           1249   | 00001)<br>Chemothin<br>Events<br>25<br>20<br>27<br>37<br>23<br>8<br>33<br>27<br>21<br>25<br>29<br>25<br>29<br>25<br>29<br>25<br>29<br>25<br>29<br>25<br>29<br>25<br>29<br>25<br>29<br>25<br>29<br>25<br>20<br>27<br>21<br>15<br>43<br>26<br>44<br>28<br>26<br>15<br>29<br>25<br>29<br>25<br>29<br>25<br>20<br>25<br>20<br>27<br>23<br>25<br>20<br>25<br>20<br>27<br>23<br>25<br>20<br>25<br>20<br>25<br>20<br>25<br>20<br>25<br>20<br>25<br>20<br>25<br>20<br>25<br>20<br>25<br>20<br>25<br>20<br>25<br>20<br>25<br>20<br>25<br>20<br>25<br>20<br>25<br>20<br>25<br>20<br>25<br>20<br>25<br>20<br>25<br>20<br>25<br>20<br>25<br>20<br>25<br>20<br>25<br>20<br>25<br>20<br>25<br>20<br>25<br>20<br>25<br>20<br>25<br>20<br>25<br>20<br>25<br>20<br>25<br>20<br>25<br>20<br>25<br>20<br>25<br>20<br>25<br>20<br>25<br>20<br>25<br>20<br>25<br>20<br>25<br>20<br>25<br>20<br>25<br>20<br>25<br>20<br>25<br>20<br>25<br>20<br>25<br>20<br>25<br>20<br>25<br>20<br>25<br>20<br>25<br>20<br>25<br>20<br>25<br>28<br>25<br>28<br>25<br>28<br>25<br>28<br>25<br>28<br>25<br>28<br>25<br>28<br>25<br>28<br>25<br>28<br>25<br>28<br>25<br>31<br>26<br>25<br>38<br>25<br>38<br>25<br>38<br>25<br>25<br>28<br>25<br>38<br>25<br>25<br>28<br>25<br>38<br>25<br>25<br>28<br>25<br>38<br>25<br>25<br>28<br>25<br>38<br>25<br>25<br>28<br>25<br>38<br>25<br>25<br>28<br>25<br>25<br>28<br>25<br>25<br>28<br>25<br>25<br>26<br>25<br>25<br>26<br>25<br>25<br>26<br>25<br>26<br>25<br>26<br>25<br>26<br>25<br>26<br>26<br>25<br>26<br>26<br>26<br>26<br>26<br>26<br>26<br>26<br>26<br>26  | Prapy           Total           31           300           44           51           27           12           54           35           311           40           390           400           63           355           43           40           63           355           43           40           63           355           43           40           55           43           40           35           255           43           40           35           26           67           40           50           50           1244  | Weight 2,5% 2,4% 3,5% 4,1% 2,2% 4,3% 2,8% 2,5% 3,3% 3,1% 3,2% 4,3% 2,8% 2,8% 2,8% 2,8% 2,8% 2,8% 2,8% 2,8  | Risk Difference<br>M-H, Fixed, 95% CI<br>0.16 [0.01, 0.31]<br>0.13 [0.09, 0.36]<br>0.00 [-0.20, 0.20]<br>0.04 [-0.13, 0.21]<br>-0.07 [-0.27, 0.14]<br>0.25 [-0.06, 0.56]<br>0.28 [0.12, 0.43]<br>0.03 [-0.16, 0.22]<br>0.01 [-0.12, 0.32]<br>0.04 [-0.21, 0.21]<br>0.01 [-0.12, 0.32]<br>0.05 [-0.13, 0.32]<br>0.05 [-0.14, 0.25]<br>0.05 [-0.09, 0.20]<br>0.19 [0.01, 0.37]<br>0.04 [-0.16, 0.24]<br>0.03 [-0.27, 0.24]<br>0.03 [-0.17, 0.23]<br>0.06 [-0.14, 0.25]<br>0.05 [-0.09, 0.20]<br>0.19 [0.01, 0.37]<br>0.04 [-0.16, 0.24]<br>0.03 [-0.17, 0.23]<br>0.06 [-0.22, 0.34]<br>0.22 [0.05, 0.40]<br>0.22 [0.05, 0.40]<br>0.20 [0.00, 0.40]<br>0.23 [0.08, 0.38]<br>0.30 [0.11, 0.49]<br>0.11 [0.08, 0.15]  | 0.01 0.1 1 10 10 Favours [Kanglaite] Favours [Chemotherapy] Risk Difference M-H, Fixed, 95% Ci          |

Figure 8 The analysis of DCR between two groups. DCR, disease control rate.

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| Table 4 S | Subgroup | analysis | of ORR | for | each | variable |
|-----------|----------|----------|--------|-----|------|----------|
|-----------|----------|----------|--------|-----|------|----------|

| Veriable                             | No of trials  | No. of participa | ants  | Freeture DD (050/ CI) | Duoluo <sup>a</sup> |
|--------------------------------------|---------------|------------------|-------|-----------------------|---------------------|
| variable                             | No. of trials | ORR: CR + PR     | Total | Fracture, RR (95% CI) | P value             |
| Kanglaite dose                       |               |                  |       |                       | 0.32                |
| 200 mL                               | 20            | 677              | 1628  | 1.37 (1.22, 1.54)     |                     |
| Other <sup>b</sup>                   | 9             | 243              | 679   | 1.55 (1.26, 1.90)     |                     |
| Type of first-line platinum-based ch | emotherapy    |                  |       |                       | 0.59                |
| GP                                   | 19            | 624              | 1538  | 1.34 (1.19, 1.52)     |                     |
| NP                                   | 5             | 153              | 374   | 1.68 (1.30, 2.18)     |                     |
| DP                                   | 4             | 131              | 348   | 1.52 (1.15, 2.01)     |                     |
| TP                                   | 3             | 77               | 268   | 1.35 (0.92, 1.98)     |                     |
| AP                                   | 1             | 18               | 49    | 1.30 (0.62, 2.73)     |                     |
| Cycle of chemotherapy                |               |                  |       |                       | 0.67                |
| 2-cycle                              | 14            | 445              | 1030  | 1.45 (1.25, 1.67)     |                     |
| Other <sup>c</sup>                   | 18            | 558              | 1547  | 1.38 (1.21, 1.58)     |                     |
| Evaluation criteria                  |               |                  |       |                       | 0.25                |
| WHO                                  | 15            | 457              | 1063  | 1.50 (1.30, 1.73)     |                     |
| RECIST                               | 17            | 546              | 1514  | 1.34 (1.17, 1.53)     |                     |

<sup>a</sup>P value: heterogeneity between subgroups; <sup>b</sup>other: includes Kanglaite doses about 60, 100 and 300 mL; <sup>c</sup>other: includes chemotherapy cycles about 1-cycle, 3-cycle, 4-cycle. ORR, objective response rate; CR, complete response; PR, partial response; GP, cisplatin or paraplatin and gemcitabine; NP, cisplatin or paraplatin and vinorelbine; TP, cisplatin or paraplatin and paclitaxel; DP, cisplatin or paraplatin and docetaxel; AP, cisplatin or paraplatin and pemetrexed.

criteria on the ORR (Table 4 and Figure S2A, B, C, D). Drug doses included 200 mL on the KLT medicine instruction (47) and other doses. Subgroup analysis indicated that both doses increased the ORR. Types of first-line platinum-based chemotherapy included cisplatin or paraplatin plus vinorelbine, paclitaxel, gemcitabine, docetaxel, or pemetrexed (NP, TP, GP, DP, and AP). Subgroup analysis demonstrated that only KLT plus GP, NP, and DP could increase the ORR. The differences between TP (P=0.12) and AP (P=0.49) were not statistically significant. The chemotherapy cycles included 2 cycles and more, and both cycles could increase the ORR. Tumor responses were evaluated using WHO or RECIST criteria. Subgroup analysis showed that KLT plus firstline platinum-based chemotherapy could increase the ORR using the WHO or RECIST criteria.

#### Subgroup analysis of nausea and vomiting

Subgroup analysis was performed to reveal the influence of different doses, types of first-line platinum-based chemotherapy, cycles of chemotherapy, evaluation criteria, and supportive treatment for nausea and vomiting (*Table 5* and *Figure S3A,B,C,D,E*). The subgroup analysis failed to find any significant differences among the subgroups on the dose of KLT, chemotherapy types, chemotherapy cycles, and supportive treatment. Furthermore, the other doses (P=0.07), other cycles (P=0.24), and unclear supportive treatment (P=0.08) were not statistically significant. Additionally, high subgroup differences regarding evaluation criteria (I2=84.8%, P=0.01) were observed.

# Subgroup analysis of leukopenia

The subgroup analysis failed to report any significant differences among subgroups on the dose of KLT, chemotherapy cycles, evaluation criteria, and supportive treatment for leukopenia (*Table 6* and *Figure S4A,B,C,D,E*). Additionally, other doses (P=0.28), other cycles (P=0.26), and unclear supportive treatment (P=0.09) were not statistically significant, but high subgroup differences regarding chemotherapy types (I2=66%, P=0.09) were noted.

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| Table 5 subgroup  | analysis of nausea | and vomiting for | each variable |
|-------------------|--------------------|------------------|---------------|
| - more e omogroup |                    |                  |               |

| Mariala                                    |              | No. of participant  | S     | - Fracture, RR (95% CI) | P value <sup>a</sup> | 12a   |
|--|--------------|---------------------|-------|-------------------------|----------------------|-------|
| variable                                   | NO. OF TRAIS | Nausea and vomiting | Total | Fracture, RR (95% CI)   | P value              | Γ     |
| Kanglaite dose                             |              |                     |       |                         | 0.97                 | 0%    |
| 200 ml                                     | 6            | 251                 | 481   | 0.58 (0.39, 0.87)       |                      |       |
| Other <sup>b</sup>                         | 2            | 20                  | 91    | 0.57 (0.31, 1.04)       |                      |       |
| Type of first-line platinum-based chemothe | erapy        |                     |       |                         | 0.40                 | 0%    |
| GP   | 8            | 239                 | 610   | 0.61 (0.43, 0.87)       |                      |       |
| ТР   | 1            | 40                  | 62    | 0.48 (0.31, 0.74)       |                      |       |
| Cycle of chemotherapy                      |              |                     |       |                         | 0.79                 | 0%    |
| 2-cycle                                    | 7            | 267                 | 505   | 0.59 (0.41, 0.83)       |                      |       |
| Other <sup>c</sup>                         | 2            | 12                  | 167   | 0.50 (0.15, 1.60)       |                      |       |
| Evaluation criteria                        |              |                     |       |                         | 0.01                 | 84.8% |
| Yes  | 4            | 187                 | 301   | 0.77 (0.61, 0.98)       |                      |       |
| Unclear                                    | 5            | 92                  | 371   | 0.45 (0.32, 0.63)       |                      |       |
| Supportive treatment                       |              |                     |       |                         | 0.78                 | 0%    |
| Yes  | 4            | 112                 | 226   | 0.59 (0.45, 0.76)       |                      |       |
| No   | 5            | 167                 | 446   | 0.53 (0.26, 1.08)       |                      |       |

<sup>a</sup>P value, I<sup>2</sup>: heterogeneity between subgroups; <sup>b</sup>other: includes Kanglaite doses about 60 and 300 mL; <sup>c</sup>other: includes chemotherapy cycles about 3-cycle and 4-cycle. GP, cisplatin or paraplatin and gemcitabine; NP, cisplatin or paraplatin and vinorelbine; TP, cisplatin or paraplatin and pacelitaxel; DP, cisplatin or paraplatin and docetaxel; AP, cisplatin or paraplatin and pemetrexed.

| Veriable                                      | No. of trials | No. of partici   | pants |                       | P value <sup>a</sup> | 12a  |
|---|---------------|------------------|-------|-----------------------|----------------------|------|
| variable                                      | NO. OF THAIS  | Leukopenia Total |       | Fracture, RR (95% CI) | P value              | I    |
| Kanglaite dose                                |               |                  |       |                       | 0.53                 | 0%   |
| 200 mL  | 7             | 309              | 625   | 0.64 (0.43, 0.94)     |                      |      |
| Other <sup>b</sup>                            | 3             | 34               | 176   | 0.34 (0.05, 2.45)     |                      |      |
| Type of first-line platinum-based chemotherap | ру            |                  |       |                       | 0.09                 | 66%  |
| GP  | 9             | 305              | 759   | 0.70 (0.53, 0.93)     |                      |      |
| Other <sup>c</sup>                            | 2             | 45               | 142   | 0.32 (0.14, 0.75)     |                      |      |
| Cycle of chemotherapy                         |               |                  |       |                       | 0.29                 | 9.3% |
| 2-cycle                                       | 7             | 280              | 529   | 0.56 (0.35, 0.90)     |                      |      |
| Other <sup>d</sup>                            | 4             | 70               | 372   | 0.79 (0.52, 1.19)     |                      |      |
| Evaluation criteria                           |               |                  |       |                       | 0.52                 | 0%   |
| Yes   | 6             | 235              | 387   | 0.67 (0.46, 0.99)     |                      |      |
| Unclear                                       | 5             | 115              | 514   | 0.55 (0.33, 0.90)     |                      |      |
| Supportive treatment                          |               |                  |       |                       | 0.32                 | 0%   |
| Yes   | 5             | 138              | 420   | 0.55 (0.35, 0.87)     |                      |      |
| No  | 6             | 212              | 481   | 0.74 (0.52, 1.05)     |                      |      |

 Table 6 Subgroup analysis of leukopenia for each variable

<sup>a</sup>P value, I<sup>2</sup>: heterogeneity between subgroups; <sup>b</sup>other: includes Kanglaite doses about 60, 100 and 300 mL; <sup>c</sup>other: includes chemotherapy types about DP and NP; <sup>d</sup>other: includes chemotherapy cycles about 3-cycle and 4-cycle. NP, cisplatin or paraplatin and vinorelbine; DP, cisplatin or paraplatin and docetaxel.





Figure 9 Publication bias analysis.

## Publication bias analysis (Figure 9)

The funnel plots were symmetric in ORR, DCR, and KPS (*Figure 9A*,*C*,*D*). Furthermore, no publication bias was observed in studies that objectively reported the results. The funnel plots were asymmetric in leukopenia (*Figure 9B*). These results indicated publication bias. Leukopenia was overestimated in one study (42).

# Summary of evidence

We used the Grading of Recommendations Assessment Development and Evaluation (GRADE) to report the quality of evidence for ORR, DCR, KPS, and adverse events. The quality of evidence was rated as high, moderate, low, and very low, and the results revealed a moderate or low overall quality (*Figure S5*).

# Discussion

As a traditional Chinese medicine injection, KLT is widely used in clinical patients with lung, gastric, and liver cancer (48,49) as an adjunct to chemotherapy for improving the curative effect. Some studies have shown that KLT can sensitize cancer cells to chemotherapy (50,51). One network meta-analysis (52) that included three traditional Chinese medicine injections showed KLT combined with NP had the greatest efficacy in ORR, ranking first with a probability of 71%.

#### Summary of the results

In our study, KLT plus first-line platinum-based chemotherapy improved the ORR and decreased the risk ratio of nausea, vomiting, and leukopenia in NSCLC. We observed significant clinical heterogeneity in adverse reactions, which is consistent with previous studies (7). However, previous studies failed to explain the longterm synergistic efficacy of this combination and did not perform subgroup analysis to clarify the observed clinical heterogeneity. Our subgroup analysis indicated these results were generally consistent regardless of the dose of KLT, chemotherapy cycles, and evaluation criteria for ORR. Regarding the type of chemotherapy, there were no significant differences between KLT combined with TP or AP. We did not reach any definite conclusions about AP due to only one relevant study being available. There were three studies in the TP subgroup and showed no obvious quality problems of ORR. Ma observed that the inhibitory effect of KLT plus TP in Lewis lung cancer cell lines was not significantly different from TP (53). Previous studies have

reported KLT plus paclitaxel demonstrated no significance compared to paclitaxel in advanced malignant thymoma (54). Aside from insufficient quantity, we assumed KLT combined with GP, NP, and DP, but not TP, could increase tumor responses. We applied the KPS scale to evaluate the QOL and observed that KLT significantly increased the KPS.

Furthermore, a meta-analysis with 31 studies showed that KLT could slightly increase the DCR. Therefore, we believe KLT plus first-line platinum-based chemotherapy, excluding the TP combination, could significantly increase clinical efficacy. The results indirectly indicate KLT may have a synergistic efficacy with first-line platinum-based chemotherapy.

First-line platinum-based chemotherapy demonstrates varying degrees of blood and gastrointestinal toxicity, and we selected the most common clinical adverse reactions, including nausea, vomiting, and leukopenia, to evaluate the role of KLT in preventing adverse reactions. For nausea and vomiting, further subgroup analysis indicated that clear evaluation criteria were the source of heterogeneity. For leukopenia, further subgroup analysis showed that variability in chemotherapy type was the source of heterogeneity, and leukopenia was overestimated in one study (41). It is worth mentioning that using use of supportive treatment during KLT plus first-line platinum-based chemotherapy would not decrease the risk ratio of both adverse reactions compared with chemotherapy alone, which has not been mentioned in previous studies (48,55). However, there are many factors in clinical settings, including supportive treatment, treatment dosage, type of chemotherapy. Hence, we assumed an indefinite conclusion regarding the role of KLT in the matter.

# Strengths and limitations

This is the most detailed meta-analysis focusing on the efficacy of ORR, KPS, and adverse reactions, and can thus better inform the clinical application of KLT. However, some limitations this study should be noted. First, only Chinese and English databases were searched, and some relevant studies might have been excluded due to language restrictions. Furthermore, all of the included studies were published in China, which might have led to publication bias. Secondly, only 17 studies reported the random allocation methods, but no study provided detailed information on the random allocation concealment.

Moreover, 14 participants withdrew from the clinical study with 5 reporting acute/subacute toxicity; this

could have influenced the outcome of adverse reactions. Additionally, three studies demonstrated selective reporting concerning acute/subacute toxicity, and 1 study did so concerning KPS. Thirdly, the methods used to classify studies as high quality might have been relatively lenient, and other researchers may have selected different definitions for study quality.

# Suggestions for future clinical trials

Differences in survival rates are of utmost importance to clinicians and NSCLC patients alike. We are similarly interested in the long-term synergistic efficacy of KLT plus first-line platinum-based chemotherapy, particularly as it relates to survival and deterioration rate effect. In our meta-analysis, three studies (19,28,45) mentioned MST, and the combined treatment showed a positive effect on MST compared with first-line platinum-based chemotherapy merely. We noticed that evidence concerning long-term synergistic efficacy, such as overall survival and progressionfree survival, was still insufficient. Apart from that, we noticed that some studies demonstrated the survival rate using a life table to show the changes vividly, but only complete reporting may bring meaningful work to NSCLC treatment. Therefore, we appeal to clinical researchers to include short-term and long-term synergistic efficacy with the specific normative data type and regard it as a vital outcome in further research.

# Conclusions

The evidence indicates KLT plus first-line platinumbased chemotherapy, except the combination with TP, may significantly improve the clinical efficacy in patients with advanced NSCLC. With supportive treatments, this combination demonstrated a lower risk of nausea, vomiting, and leukopenia, and positively affected MST and KPS. These results indicate KLT may indirectly have ameliorative and synergistic efficacy with first-line platinum-based chemotherapy. Finally, many shortcomings in clinical trial methodology resulted in an inadequate assessment of clinical efficacy and safety. We are thus eager to evaluate larger-scale RCTs or real-world studies to present an indepth review in the near future.

## **Acknowledgments**

We would like to thank Session of Evidence-based Clinical

Club (EBC) for the direction on the manuscripts.

*Funding:* This work was financially supported by National key research and development plan of China (No. 2018YFC1707405) and National Natural Science Foundation of China (No. 81774289, 81473463). The funders had no role in study design, data collection and analysis, decision to publish, or preparation of the manuscript.

# Footnote

Reporting Checklist: The authors have completed the PRISMA 2009 reporting checklist (available at http://dx.doi.org/10.21037/apm-20-616)

Data Sharing Statement: Available at http://dx.doi. org/10.21037/apm-20-616

Peer Review File: Available at http://dx.doi.org/10.21037/ apm-20-616

*Conflicts of Interest:* All authors have completed the ICMJE uniform disclosure form (available at http://dx.doi. org/10.21037/apm-20-616). The authors have no conflicts of interest to declare.

*Ethical Statement:* The authors are accountable for all aspects of the work in ensuring that questions related to the accuracy or integrity of any part of the work are appropriately investigated and resolved. As a systematic review and meta-analysis, Ethical approval was not required as materials of this study had been published.

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**Cite this article as:** Li J, Li HZ, Zhu GH, Gao RK, Zhang Y, Hou W, Li J. Efficacy and safety of Kanglaite injection combined with first-line platinum-based chemotherapy in patients with advanced NSCLC: a systematic review and meta-analysis of 32 RCTs. Ann Palliat Med 2020;9(4):1518-1535. doi: 10.21037/apm-20-616

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| Search     | Add to builder | Query   | Items found    | Time     |
|------------|----------------|---|----------------|----------|
| <u>#23</u> | Add            | Search ((([Kanglaite) OR Coix Seed Oil)) AND ((non-small cell lung carcinoma[Title/Abstract]) OR<br>NSCLC[Title/Abstract])) AND ((((((([Kandomized Controled Trials) OR Randomized Controled Trial)<br>OR controlled clinical trial) OR placebo) OR drug therapy) OR randomly) OR trial) OR groups)) NOT<br>((animals[MeSH Terms]) NOT humans[MeSH Terms])) | 4              | 06:15:09 |
| #22        | Add            | Search (Kanglaite) OR Coix Seed Oil   | <u>70</u>      | 06:14:52 |
| <u>#21</u> | Add            | Search ((((((Randomized Controled Trials) OR Randomized Controled Trial) OR controlled clinical<br>trial) OR placebo) OR drug therapy) OR randomly) OR trial) OR groups)) NOT ((animals[MeSH Terms])<br>NOT humans[MeSH Terms])   | 4802789        | 06:14:16 |
| #20        | Add            | Search (animals[MeSH Terms]) NOT humans[MeSH Terms]   | 4632487        | 06:14:07 |
| <u>#19</u> | Add            | Search ((((((Randomized Controled Trials) OR Randomized Controled Trial) OR controlled clinical<br>trial) OR placebo) OR drug therapy) OR randomly) OR trial) OR groups   | <u>5618401</u> | 06:13:14 |
| #18        | Add            | Search groups   | 2094991        | 06:12:30 |
| <u>#17</u> | Add            | Search trial  | 1354201        | 06:12:24 |
| #16        | Add            | Search randomly   | 320848         | 06:12:18 |
| #15        | Add            | Search drug therapy   | 3111089        | 06:12:07 |
| #14        | Add            | Search placebo  | 222394         | 06:11:58 |
| #13        | Add            | Search controlled clinical trial  | 713222         | 06:11:48 |
| #12        | Add            | Search Randomized Controled Trial   | 32             | 06:08:42 |
| #11        | Add            | Search Randomized Controled Trials  | 16             | 06:08:33 |
| #9         | Add            | Search Coix Seed Oil  | 26             | 06:07:57 |
| #8         | Add            | Search (non-small cell lung carcinoma[Title/Abstract]) OR NSCLC[Title/Abstract]   | 41561          | 06:07:35 |
| #7         | Add            | Search Kanglaite  | 46             | 06:06:55 |

Figure S1 Electronic search strategy for PubMed.

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|                        | Kangla     | iite    | Contr       | lor     |          | Risk Ratio         | Risk Ratio  |
|------------------------|------------|---------|-------------|---------|----------|--------------------|---|
| Study or Subgroup      | Events     | Total   | Events      | Total   | Weight   | M-H, Fixed, 95% Cl | M-H, Fixed, 95% Cl  |
| 1.4.1 200 ml           |            |         |             |         |          |                    |   |
| Bao H 2019             | 25         | 31      | 17          | 31      | 4.5%     | 1.47 [1.02, 2.11]  |   |
| Chen C 2018            | 16         | 30      | 11          | 30      | 2.9%     | 1.45 [0.82, 2.59]  |   |
| Chen Y 2018            | 22         | 51      | 18          | 51      | 4.7%     | 1.22 [0.75, 1.99]  |   |
| chen YZ 2003           | 13         | 28      | 9           | 27      | 2.4%     | 1.39 [0.72, 2.71]  |   |
| luang ZB 2010          | 19         | 35      | 13          | 35      | 3.4%     | 1.46 [0.86, 2.48]  |   |
| a JN 2018              | 15         | 31      | 10          | 31      | 2.6%     | 1.50 [0.80, 2.81]  | and the second se |
| i XD 2016              | 24         | 39      | 14          | 39      | 3.7%     | 1.71 [1.05, 2.79]  | · · · · · · · · · · · · · · · · · · ·   |
| iu F 2019              | 40         | 63      | 26          | 63      | 6.8%     | 1.54 [1.09, 2.18]  |   |
| iu JQ 2011             | 15         | 35      | 12          | 35      | 3.2%     | 1.25 (0.69, 2.27)  |   |
| iu XH 2016             | 27         | 55      | 17          | 55      | 4.5%     | 1.59 (0.98. 2.56)  |   |
| iu Y 2015              | 18         | 43      | 14          | 43      | 3.7%     | 1.29 (0.74, 2.24)  |   |
| ong SG 2017            | 17         | 42      | 15          | 40      | 4.0%     | 1 08 10 63 1 861   |   |
| U Q 2018               | 24         | 47      | 14          | 47      | 37%      | 1 71 (1 02 2 89)   |   |
| un SQ 2012             | 15         | 35      | 11          | 35      | 2 9%     | 1 36 (0 73 2 54)   |   |
| and HY 2016            | 10         | 24      | 8           | 25      | 21%      | 1 30 10 62 2 731   |   |
| Vang 1 2014            | 10         | 43      | 12          | 43      | 2.7%     | 1 59 10 99 2 951   |   |
| ang C 2014             | 22         | 40      | 21          | 40      | 5.5%     | 1 10 10 71 1 701   |   |
| ana Q12002             | 20         | 25      | 7           | 26      | 1 0%     | 1 24 10 59 2 041   |   |
| ang 55 2005            | 10         | 70      | 16          | 67      | 1.0%     | 1.34 [0.33, 3.04]  |   |
| a CV 2010              | 22         | 40      | 21          | 40      | 5.5%     | 1.05 10 70 1 571   |   |
| e Cr 2015              | 44         | 916     | 41          | 812     | 75.0%    | 1 37 [1 22 1 54]   | ▲   |
| into lovente           | 202        | 010     | 205         | 012     | 1.0.0 10 | inti [nee, mail    |   |
| lotaregeneity Chiz-    | 6 02 df-   | 10 /0   | 1 000/13    | - 00    |          |                    |   |
| reterogeneity, chir=   | 0.03, ui = | 13(1-   | = 1.00), P  | = 0 %   |          |                    |   |
| est for overall ellect | 2= 0.32 (  | P = 0.0 | 0001)       |         |          |                    |   |
| A 2 Other              |            |         |             |         |          |                    |   |
| Han VO 2000            | 6          | 12      |             | 12      | 1.100    | 1 36 10 44 3 661   |   |
| uan XQ 2009            | 22         | 12      | 4           | 12      | 1.1%     | 1.25 [0.44, 3.55]  |   |
| 0 1 2017               | 22         | 34      |             | 04      | 2.970    | 2.00 [1.08, 3.71]  |   |
| L 2012                 | 22         | 30      | 9           | 40      | 2.370    | 0.34 [0.40, 2.17]  |   |
| lang J 2018            | 23         | 40      | 10          | 40      | 2.0%     | 2.30 [1.26, 4.19]  |   |
| lang SG 2014           | 10         | 23      | 8           | 20      | 2.2%     | 1.09 [0.53, 2.21]  |   |
| rang CY 2018           | 18         | 42      | 17          | 42      | 4.5%     | 1.06 [0.64, 1.76]  |   |
| vang Y 2017            | 26         | 36      | 12          | 36      | 3.2%     | 2.17 [1.31, 3.59]  | 1   |
| ang L 2016             | 13         | 35      | 10          | 35      | 2.6%     | 1.30 [0.66, 2.56]  |   |
| u 1 2015               | 23         | 60      | 14          | 60      | 3.7%     | 1.64 [0.94, 2.88]  |   |
| ubtotal (95% Cl)       |            | 340     | 10          | 339     | 25.0%    | 1.55 [1.26, 1.90]  | <b>T</b>  |
| otal events            | 148        |         | 95          |         |          |                    | 1.0 million   |
| leterogeneity. Chi* =  | 8.99, df=  | 8 (P =  | 0.34); 1ª = | = 11%   |          |                    |   |
| est for overall effect | Z= 4.12 (  | P < 0.0 | 001)        |         |          |                    |   |
| otal (95% CI)          |            | 1156    |             | 1151    | 100.0%   | 1.41 (1.28, 1.57)  | •   |
| intal evente           | 640        | 1100    | 380         |         |          | The Lord 1991      |   |
| lataraganaity Chiz-    | 16 71 4    | - 20 /0 | -0.06       | 17 - 04 |          |                    | 1   |
| eterogeneny. Chi*=     | 7-6.70     | - 20 (P | - 0.99),    |         |          |                    | 0.01 0.1 1 10 10  |
| est for overall effect | 2= 0.70 (  | P < 0.0 | 0001)       |         |          | 0.00               | Favours [experimental] Favours [control]  |

**Risk Ratio** 

1.47 [1.02, 2.11]

1.39 [0.72, 2.71]

1.25 [0.44, 3.55]

1.46 [0.86, 2.48]

1.50 [0.80, 2.81] 1.71 [1.05, 2.79]

2.30 [1.26, 4.19] 1.09 [0.53, 2.21]

1.54 [1.09, 2.18] 1.25 [0.69, 2.27]

1.71 [1.02, 2.89]

1.30 [0.62, 2.73]

1.21 [0.67, 2.18]

1.05 [0.70, 1.57]

1.45 [1.25, 1.67]

1.45 [0.82, 2.59]

1.13 [0.65, 1.97] 1.22 [0.75, 1.99]

2.00 [1.08, 3.71]

1.14 [0.65, 2.02] 0.94 [0.40, 2.17]

1.59 [0.98, 2.56] 1.29 [0.74, 2.24]

1.08 [0.63, 1.86] 1.36 [0.73, 2.54]

1.06 [0.64, 1.76]

1.58 [0.88, 2.85] 2.17 [1.31, 3.59]

1.10 [0.71, 1.70] 1.30 [0.66, 2.56]

1.34 [0.59, 3.04] 1.64 [0.94, 2.88]

2.50 [1.06, 5.92] 1.38 [1.21, 1.58]

1.41 [1.28, 1.56]

0.01

0.1

Favours [exper

**Risk Ratio** 

M-H. Fixed, 95% Cl

4444444444444

Kanglaite

Total events 264 181 Heterogeneity: Chi<sup>2</sup> = 7.09, df = 13 (P = 0.90); l<sup>2</sup> = 0% Test for overall effect: Z = 5.09 (P < 0.00001)

324

588

Heterogeneity: Chi\* = 19.51, df = 31 (P = 0.95); I\* = 0%

Test for overall effect: Z = 6.91 (P < 0.00001)

Heterogeneity: Chi# = 12.20, df = 17 (P = 0.79); I# = 0% Test for overall effect: Z = 4.75 (P < 0.00001)

31

40 518

30

1291

Test for subaroup differences: Chi<sup>2</sup> = 0.19, df = 1 (P = 0.67), l<sup>2</sup> = 0%

С

Study or Subgroup

1.5.1 2-cycle Bao H 2019

Chen YZ 2003

Guan XQ 2009

Huang ZB 2010

**Jia JN 2018** LI XD 2016 Liang J 2018 Liang SG 2014

Liu F 2019 Liu JQ 2011

Mu Q 2018

Yao J 2017

Ye CY 2019

1.5.2 other

Chen C 2018

Chen W 2018

Chen Y 2018

He LT 2017

LI HY 2017 LI L 2012

Liu XH 2016 Liu Y 2015

Long SG 2017

Wang CY 2018

Sun SQ 2012

Wang L 2014 Wang Y 2017

Yan QH 2018 Yang L 2016

Yang SJ 2003 Yu T 2015

Total events

Total (95% CI)

Total events

Zhang MM 2019 Subtotal (95% CI)

Wang HY 2016

Subtotal (95% CI)

Control

17

9

4

13

10 14 10

8

26 12 14

8 15 21

11 15

18 11

14 41 40 9

7 14 6

234

415

Events Total Events Total Weight M-H, Fixed, 95% Cl

25 67

40

512

30

44 51 54

65 43

1286 100.0%

4.1%

2.2%

3.1%

2.4% 3.4%

2.4% 2.1%

6.3% 2.9%

3.4%

3.7%

5.0% 43.7%

2.6%

3.6% 4.3% 2.6%

3.4% 2.1%

4.1% 3.4%

3.7%

4.1% 2.9%

2.9% 5.0% 2.4%

1.7% 3.4%

1.4% 56.3%

| В | Study or Subgroup  | Kanglaite Chemotherapy<br>Events Total Events Total |                               |                                  | Weight  | Risk Ratio<br>M-H, Fixed, 95% Cl | Risk Ratio<br>M-H, Fixed, 95% Cl |   |  |
|---|--|---|-------------------------------|----------------------------------|---------|----------------------------------|----------------------------------|---|--|
|   | 1.1.1 GP<br>Bao H 2019                                       | 25  | 31                            | 17                               | 31      | 4.1%                             | 1 47 (1 02 2 11)                 |   |  |
|   | Chen C 2018  | 16  | 30                            | 11                               | 30      | 2.6%                             | 1.45 [0.82, 2.59]                |   |  |
|   | Chen W 2016  | 17  | 44                            | 15                               | 44      | 3.6%                             | 1.13 [0.65, 1.97]                |   |  |
|   | Guan XQ 2009   | 5   | 12                            | 4                                | 12      | 1.0%                             | 1.25 [0.44, 3.55]                |   |  |
|   | Huang ZB 2010  | 19  | 35                            | 13                               | 35      | 3.1%                             | 1.46 [0.86, 2.48]                |   |  |
|   | Li HY 2017   | 16  | 41                            | 14                               | 41      | 3.4%                             | 1.14 [0.65, 2.02]                |   |  |
|   | Liu F 2019   | 40  | 63                            | 26                               | 63      | 6.3%                             | 1.54 [1.09, 2.18]                |   |  |
|   | Liu JQ 2011  | 15  | 35                            | 12                               | 35      | 2.9%                             | 1.25 (0.69, 2.27)                |   |  |
|   | Liu Y 2015   | 18  | 43                            | 14                               | 43      | 3.4%                             | 1.29 [0.74, 2.24]                |   |  |
|   | Sun SQ 2012  | 15  | 35                            | 11                               | 35      | 2.6%                             | 1.36 [0.73, 2.54]                |   |  |
|   | Wang L 2014  | 19  | 43                            | 12                               | 43      | 2.9%                             | 1.58 [0.88, 2.85]                |   |  |
|   | Wang Y 2017<br>Yan QH 2018                                   | 26  | 36                            | 12                               | 36      | 2.9%                             | 2.17 [1.31, 3.59]                |   |  |
|   | Yao J 2017   | 19  | 70                            | 15                               | 67      | 3.7%                             | 1.21 (0.67, 2.18)                |   |  |
|   | Ye CY 2019   | 22  | 40                            | 21                               | 40      | 5.0%                             | 1.05 [0.70, 1.57]                |   |  |
|   | Zhang MM 2019<br>Subtotal (95% CI)                           | 15  | 50                            | 6                                | 765     | 1.4%                             | 2.50 [1.06, 5.92]                | •   |  |
|   | Total events<br>Heterogeneity: Chi <sup>2</sup> =            | 359<br>11.00, df=                                   | 18 (P :                       | 265<br>= 0.89); I <sup>2</sup> = | 0%      |                                  | and a set                        |   |  |
|   | Test for overall effect                                      | Z= 4.78 (F  | e = 0.00                      | 1001)                            |         |                                  |                                  |   |  |
|   | He LT 2017   | 22  | 54                            | 11                               | 54      | 2.6%                             | 2.00 [1.08, 3.71]                |   |  |
|   | Jia JN 2018  | 15  | 31                            | 10                               | 31      | 2.4%                             | 1.50 [0.80, 2.81]                |   |  |
|   | Wang CY 2018   | 18  | 47                            | 17                               | 42      | 4.1%                             | 1.06 [0.64, 1.76]                |   |  |
|   | Subtotal (95% CI)  |   | 174                           |                                  | 174     | 12.5%                            | 1.52 [1.15, 2.01]                | •   |  |
|   | Total events<br>Heterogeneity: Chi <sup>z</sup> =            | 79<br>2.92, df = 3                                  | (P=0                          | 52<br>.40); I <sup>2</sup> = 0   | 96      |                                  |                                  |   |  |
|   | Test for overall effect                                      | Z= 2.93 (F  | r = 0.00                      | (3)                              |         |                                  |                                  |   |  |
|   | Chen YZ 2003   | 13  | 28                            | 9                                | 27      | 2.2%                             | 1.39 (0.72, 2.71)                |   |  |
|   | Liang J 2018   | 24  | 39                            | 14                               | 39      | 3.4%                             | 2.30 (1.26 4 19)                 |   |  |
|   | Liu XH 2016  | 27  | 55                            | 17                               | 55      | 4.1%                             | 1.59 [0.98, 2.56]                |   |  |
|   | Yang SJ 2003<br>Sublotal (05% CD                             | 9   | 25                            | 7                                | 26      | 1.7%                             | 1.34 [0.59, 3.04]                |   |  |
|   | Total events   | 96  | 10/                           | 57                               | 187     | 13.7%                            | 1.00 [1.30, 2.18]                | The second se |  |
|   | Heterogeneity: Chi <sup>2</sup> =<br>Test for overall effect | 1.72, df = 4<br>Z = 3.97 (F                         | (P = 0<br>< 0.00              | (79); (*= 0<br>(01)              | %       |                                  |                                  |   |  |
|   | 1.1.4 TP   |   | 4                             |                                  | - 32    |                                  |                                  |   |  |
|   | Yang L 2018  | 8   | 38                            | 9                                | 40      | 2.1%                             | 0.94 [0.40, 2.17]                |   |  |
|   | Yu T 2015  | 23  | 60                            | 14                               | 60      | 3.4%                             | 1.64 [0.94, 2.88]                |   |  |
|   | Subtotal (95% CI)  | 1.1   | 133                           |                                  | 135     | 7,9%                             | 1.35 [0.92, 1.98]                | •   |  |
|   | Total events<br>Heterogeneity: Chi <sup>2</sup> =            | 44<br>1.21, df = 1                                  | (P=0                          | 33<br>.55); I⁼ = 0               | 96      |                                  |                                  |   |  |
| đ | 1.1.5 AP   | 2-1.03 (P   | - 0.12                        | /                                |         |                                  |                                  |   |  |
|   | Wang HY 2016   | 10  | 24                            | 8                                | 25      | 1.9%                             | 1.30 [0.62, 2.73]                |   |  |
|   | Subtotal (95% CI)  |   | 24                            |                                  | 25      | 1.9%                             | 1.30 [0.62, 2.73]                |   |  |
|   | Heterogeneity: Not an  | oplicable   |                               | 8                                |         |                                  |                                  |   |  |
|   | Test for overall effect                                      | Z= 0.70 (F  | = 0.49                        | 9                                |         |                                  |                                  |   |  |
|   | Total (95% CI)   | 1   | 1291                          |                                  | 1286    | 100.0%                           | 1.41 [1.28, 1.56]                | •   |  |
|   | Total events<br>Heterogeneity: Chi <sup>2</sup> =            | 588<br>19,51, df=                                   | 31 (P =                       | 415<br>= 0.95): I*=              | : 0%    |                                  |                                  | ta di time  |  |
|   | Test for overall effect<br>Test for subgroup dif             | Z = 6.91 (F<br>ferences: C                          | < 0.00<br>hi <sup>2</sup> = 2 | 1001)<br>81. df= 4 i             | P = 0.5 | 9), P= 0%                        |                                  | 0.01 0.1 1 10 10<br>Favours (Chemotherapy) Favours (Kanglaite)  |  |
| D | Study or Subaroun  | Kangl   | aite                          | Contr                            | Total   | Weight                           | Risk Ratio                       | Risk Ratio<br>M-H. Fixed, 95% Cl  |  |
|   | 1.6.1 WHO<br>Bao H 2019                                      | 25  | 21                            | 17                               | 31      | 4 1 96                           | 1.47 11.02 2.111                 |   |  |
|   | Chen Y 2018  | 22  | 51                            | 18                               | 51      | 4.3%                             | 1.22 [0.75, 1.99]                |   |  |
|   | Chen YZ 2003   | 13  | 28                            | 9                                | 27      | 2.2%                             | 1.39 [0.72, 2.71]                |   |  |
|   | Guan XQ 2009   | 5   | 12                            | 4                                | 12      | 1.0%                             | 1.25 [0.44, 3.55]                |   |  |
|   | Jia JN 2018  | 19  | 35                            | 13                               | 35      | 2.4%                             | 1.50 [0.80, 2.48]                |   |  |
|   | Li XD 2016   | 24  | 39                            | 14                               | 39      | 3.4%                             | 1.71 [1.05, 2.79]                |   |  |
|   | Liang J 2018   | 23  | 40                            | 10                               | 40      | 2.4%                             | 2.30 [1.26, 4.19]                | 1.00  |  |
|   | Liang SG 2014  | 10  | 23                            | 8                                | 20      | 2.1%                             | 1.09 [0.53, 2.21]                |   |  |
|   | Sun SQ 2012  | 15  | 35                            | 11                               | 35      | 2.6%                             | 1.36 [0.73, 2.54]                |   |  |
|   | Wang Y 2017  | 26  | 36                            | 12                               | 36      | 2.9%                             | 2.17 [1.31, 3.59]                |   |  |
|   | Yang SJ 2003   | 9   | 25                            | 7                                | 26      | 1.7%                             | 1.34 [0.59, 3.04]                |   |  |
|   | Yu T 2015  | 22  | 40                            | 21                               | 40      | 3.4%                             | 1.05 [0.70, 1.57]                |   |  |
|   | Subtotal (95% CI)  | 23  | 533                           | 14                               | 530     | 43.9%                            | 1.50 [1.30, 1.73]                | ◆ a   |  |
|   | Total events<br>Heterogeneity: Chi <sup>2</sup>              | 275<br>= 9.46, df =                                 | 14 (P                         | 182<br>= 0.80); P                | = 0%    |                                  | 11-03                            |   |  |
|   | Test for overall effec                                       | t Z = 5.62  | (P < 0,                       | 00001)                           |         |                                  |                                  |   |  |
|   | Chen C 2018  | 16  | 30                            | 11                               | 30      | 2.6%                             | 1.45 [0.82, 2.59]                |   |  |
|   | Chen W 2016  | 17  | 44                            | 15                               | 44      | 3.6%                             | 1.13 [0.65, 1.97]                |   |  |
|   | He LT 2017   | 22  | 54                            | 11                               | 54      | 2.6%                             | 2.00 [1.08, 3.71]                |   |  |
|   | LI L 2012  | 8   | 38                            | 9                                | 40      | 2.1%                             | 0.94 [0.40. 2.17]                |   |  |
|   | Liu F 2019   | 40  | 63                            | 26                               | 63      | 6.3%                             | 1.54 [1.09, 2.18]                |   |  |
|   | Liu JQ 2011  | 15  | 35                            | 12                               | 35      | 2.9%                             | 1.25 [0.69, 2.27]                |   |  |
|   | Liu Y 2015   | 19  | 43                            | 14                               | 43      | 3.4%                             | 1,29 [0.98, 2,56]                |   |  |
|   | Long SG 2017   | 17  | 42                            | 15                               | 40      | 3.7%                             | 1.08 [0.63, 1.86]                |   |  |
|   | Wang CY 2018   | 18  | 42                            | 17                               | 42      | 4.1%                             | 1.06 [0.64, 1.76]                |   |  |
|   | Wang HY 2016   | 10  | 24                            | 8                                | 25      | 1.9%                             | 1.30 [0.62, 2.73]                |   |  |
|   | Yan QH 2014  | 19  | 43                            | 21                               | 43      | 5.0%                             | 1.08 [0.88, 2.85]                | -   |  |
|   | Yang L 2016  | 13  | 35                            | 10                               | 35      | 2.4%                             | 1.30 [0.66, 2.56]                |   |  |
|   | Yao J 2017   | 19  | 70                            | 15                               | 67      | 3.7%                             | 1.21 [0.67, 2.18]                |   |  |
|   | Zhang MM 2019<br>Subtotal (95% Ch                            | 15  | 50                            | 6                                | 50      | 1.4%                             | 2.50 [1.06, 5.92]                | •   |  |
|   | Total events   | 313   |                               | 233                              |         |                                  | in the second                    |   |  |
|   | Heterogeneity: Chi*<br>Test for overall effect               | = 8.90, df =<br>1 Z = 4.26                          | 16 (P<br>(P < 0.              | = 0.92); /²<br>0001)             | = 0%    |                                  |                                  |   |  |
|   |  |   |                               |                                  |         |                                  |                                  |   |  |

Figure S2 Subgroup analysis of ORR for each variable. (A) Subgroup analysis of ORR for Kanglaite dose; (B) subgroup analysis of ORR for type of first-line platinum-based chemotherapy; (C) subgroup analysis of ORR for cycle of chemotherapy; (D) subgroup analysis of ORR for evaluation criteria. ORR, objective response rate.

100

10

mental] Favours [control]

Total events

588

Test for subaroup differences: ChI<sup>2</sup> = 1.31. df = 1 (P = 0.25). I<sup>2</sup> = 23.7%

Heterogeneity: Chi# = 19.51, df = 31 (P = 0.95); I# = 0%

Test for overall effect: Z = 6.91 (P < 0.00001)

415

0.01

0.1

Favours [exper

100

10

Favours (control)

Chemotherapy Risk Ratio Kanglaite **Risk Ratio** Total Weight M-H, Random, 95% Cl Study or Subaroup **Events Total** Events M-H, Random, 95% CI 2.8.1 200ml Bao H 2019 0.29 (0.06, 1.27) 4.4% 2 31 31 Jia JN 2018 Liu JQ 2011 17.4% 0.48 [0.31, 0.74] 0.71 [0.47, 1.07] 13 17 31 35 27 31 35 24 0.50 [0.19, 1.31] 0.88 [0.76, 1.02] Wang L 2014 5 35 10 35 8.4% Yao J 2017 55 70 60 67 22.9% Ye CY 2019 9 40 22 40 13.2% 0.41 [0.22, 0.78] Subtotal (95% CI) 242 239 84.2% 0.58 [0.39, 0.87] 101 150 Total events Heterogeneity: Tau<sup>2</sup> = 0.16; Chi<sup>2</sup> = 20.24, df = 5 (P = 0.001); I<sup>2</sup> = 75% Test for overall effect: Z = 2.66 (P = 0.008) 2.8.2 other Guan XQ 2009 6 12 10 12 13.6% 0.60 [0.32, 1.12] Wang Y 2017 1 35 3 32 2.2% 0.30 (0.03, 2.78) 15.8% Subtotal (95% CI) 47 44 0.57 [0.31, 1.04] 13 Total events Heterogeneity: Tau<sup>2</sup> = 0.00; Chi<sup>2</sup> = 0.40, df = 1 (P = 0.53); l<sup>2</sup> = 0% Test for overall effect: Z = 1.84 (P = 0.07) Total (95% CI) 289 0.58 [0.41, 0.82] 283 100.0% Total events 108 163 Heterogeneity: Tau<sup>2</sup> = 0.13; Chi<sup>2</sup> = 22.04, df = 7 (P = 0.003); i<sup>2</sup> = 68% 0.01 10 8.1 100 Test for overall effect Z = 3,08 (P = 0,002) Test for subgroup differences: ChP= 0,002 H = 1 (P = 0.97). P = 0% Favours [Kanglaite] Favours (Chemotherapy) **Risk Ratio Risk Ratio** Kanglaite Chemotherapy В Study or Subgroup **Events Total Events** Total Weight M-H, Random, 95% CI M-H, Random, 95% CI 2.10.1 GP Bao H 2019 2 31 31 4.1% 0.29 [0.06, 1.27] 0.60 [0.32, 1.12] 0.71 [0.47, 1.07] Guan XQ 2009 12 12.8% 8 10 Liu JQ 2011 17 17.2% 35 24 35 Wang L 2014 Wang Y 2017 35 35 7.8% 2.0% 0.50 [0.19, 1.31] 0.30 [0.03, 2.78] 5 10 35 32 Yao J 2017 55 70 60 67 22.4% 0.88 [0.76, 1.02] Ye CY 2019 Zhang MM 2019 9 40 22 40 12.5% 0.41 [0.22, 0.78] 50 4.6% 3 50 0.60 [0.15, 2.38] 0.61 [0.43, 0.87] 83.4% Subtotal (95% CI) 308 302 98 141 Total events Heterogeneity: Tau2 = 0.11; Chi2 = 16.45, df = 7 (P = 0.02); P = 57% Test for overall effect: Z = 2.77 (P = 0.006) 2.10.2 other Jia JN 2018 Subtotal (95% CI) 0.48 [0.31, 0.74] 0.48 [0.31, 0.74] 31 31 13 27 31 16.6% 31 16.6% 13 Total events 27 Heterogeneity: Not applicable Test for overall effect: Z = 3.29 (P = 0.001) Total (95% CI) 339 333 100.0% 0.58 [0.42, 0.81] 168 Total events 111 Heterogeneity: Tau<sup>a</sup> = 0.12; Chi<sup>a</sup> = 22.24, df = 8 (P = 0.004); l<sup>a</sup> = 64%. Test for overall effect: Z = 3.20 (P = 0.001) 0.01 0.1 10 100 Favours [Kanglaite] Favours [Chemotherapy] Test for subaroup differences: Chi<sup>2</sup> = 0.70. df = 1 (P = 0.40). l<sup>2</sup> = 0% **Risk Ratio Risk Ratio** Kanglaite Chemotherapy С Total M-H. Random, 95% Cl Study or Subgroup Events Total Events Weight M-H, Random, 95% Cl 2.9.1.2-cvcle Bao H 2019 2 4.1% 0.29 [0.06, 1.27] 31 31 10 8 12 12.8% 0.60 [0.32, 1.12] Guan XQ 2009 12 16.6% 0.48 [0.31, 0.74] 0.71 [0.47, 1.07] **Jia JN 2018** 13 31 35 27 31 35 Liu JQ 2011 17 24 Wang L 2014 5 35 10 35 7.8% 0.50 (0.19, 1.31) Yao J 2017 55 70 40 60 22 67 22.4% 0.88 [0.76, 1.02] Ye CY 2019 40 0.41 [0.22, 0.78] 9 12.5% Subtotal (95% CI) 254 251 93.4% 0.59 [0.41, 0.83] 107 160 Total events Heterogeneity: Tau# = 0.13; Chi# = 20 76, df = 6 (P = 0.002); I# = 71 % Test for overall effect: Z = 2.97 (P = 0.003) 2.9.2 other 35 50 85 Wang Y 2017 32 2.0% 0.30 (0.03, 2.78) Zhang MM 2019 3 5 50 4.8% 0.60 [0.15, 2.38] 0.50 [0.15, 1.60] Subtotal (95% CI) 82 6.6% Total events Heterogeneity: Tau<sup>2</sup> = 0.00; Chi<sup>2</sup> = 0.26, df = 1 (P = 0.61); i<sup>2</sup> = 0% Test for overall effect: Z = 1.17 (P = 0.24) Total (95% CI) 339 333 100.0% 0.58 [0.42, 0.81] Total events 111 168 Heterogeneity: Tau\* = 0.12; Chi\* = 22.24, df = 8 (P = 0.004); i\* = 64\% 0.01 0.1 10 100 Test for overall effect: Z = 3.20 (P = 0.001) Test for subgroup differences:  $Ch^{\mu}$  = 0.07. df = 1 (P = 0.79).  $l^{\mu}$  = 0% Favours [Kanglaite] Favours [Chemotherapy] Kanglaite Chemotherapy **Risk Ratio Risk Ratio** D Study or Subgroup **Events Total Events** Total Weight M-H, Random, 95% CI M-H, Random, 95% CI 2.12.1 yes Guan XQ 2009 6 12.8% 0.60 (0.32, 1.12) 12 10 12 Liu JQ 2011 Wang L 2014 35 35 35 35 17.2% 0.71 [0.47, 1.07] 0.50 [0.19, 1.31] 24 10 17 5 Yao J 2017 55 70 60 67 22.4% 0.8810.76.1.02 Subtotal (95% CI) 152 60.2% 0.77 [0.61, 0.98] 149 83 104 Total events Heterogeneity: Tau<sup>2</sup> = 0.02; Chi<sup>2</sup> = 4.30, df = 3 (P = 0.23); I<sup>2</sup> = 30% Test for overall effect: Z = 2.13 (P = 0.03) 2.12.2 unclear Bao H 2019 0.29 [0.06, 1.27] 4.1% 31 31 lia. IN 2018 16.6% 2.0% 0.48 [0.31, 0.74] 0.30 [0.03, 2.78] 13 31 35 27 31 Wang Y 2017 3 32 Ye CY 2019 12.5% 9 40 22 40 0.41 [0.22, 0.78] Zhang MM 2019 Subtotal (95% CI) 3 50 5 50 4.8% 0 60 10 15 2 38 0.45 [0.32, 0.63] 39.8% 187 184 64 Total events 28 Heterogeneity: Tau<sup>2</sup> = 0.00; Chi<sup>2</sup> = 0.85, df = 4 (P = 0.93); I<sup>2</sup> = 0% Test for overall effect: Z = 4.67 (P < 0.00001) Total (95% CI) 339 333 100.0% 0.58 [0.42, 0.81] Total events 111 168 Total events 1011 Total events 100 Heterogeneity: Tau<sup>2</sup> = 0.12; Chi<sup>2</sup> = 22.24, df = 8 (P = 0.004); l<sup>2</sup> = 64% Test for overall effect: Z = 3.20 (P = 0.001) Test for subaroup differences: Chi<sup>2</sup> = 6.57. df = 1 (P = 0.01). l<sup>2</sup> = 84.8% 0.01 0.1 10 100 Favours [Kanglaite] Favours [Chemotherapy] **Risk Ratio** Kanglaite Chemotherapy **Risk Ratio** Е Study or Subgroup Events Total Events Total Weight M-H. Random, 95% Cl M-H, Random, 95% Cl



Figure S3 Subgroup analysis of nausea and vomiting for each variable. (A) Subgroup analysis of nausea and vomiting for Kanglaite dose; (B) subgroup analysis of nausea and vomiting for type of first-line platinum-based chemotherapy; (C) subgroup analysis of nausea and vomiting for cycle of chemotherapy; (D) subgroup analysis of nausea and vomiting for evaluation criteria; (E) subgroup analysis of nausea and vomiting for supportive treatment.

Kanglaite Chemotherapy **Risk Ratio Risk Ratio** А Events Total Total Weight M-H, Random, 95% Cl Study or Subgroup Events M-H, Random, 95% CI 2.5.1 200 ml Chen Y 2018 0.83 [0.47, 1.47] 15 11.4% 51 18 51 35 31 13.4% 11.8% 0.62 [0.41, 0.93] 0.38 [0.23, 0.66] Huang ZB 2010 16 10 35 31 26 26 Jia JN 2018 Liu. IQ 2011 87 35 23 35 10.3% 0.35 [0.18, 0.67] 0.33 [0.18, 0.87] 0.70 [0.29, 1.67] 0.92 [0.47, 1.82] 43 49 8.0% 10.1% Wang L 2014 10 43 Yan QH 2018 12 13 49 70 314 Yao J 2017 60 65 67 16.4% 0 88 10 80, 0 981 Subtotal (95% CI) 311 81.4% 0.64 [0.43, 0.94] 181 128 Total events Heterogeneity:  $Tau^2 = 0.19$ ;  $Chi^2 = 30.84$ , df = 6 (P Test for overall effect: Z = 2.27 (P = 0.02) < 0.0001); |= 81% 2.5.2 other Guan XQ 2009 9 12 11 12 13.9% 0.82 [0.57, 1.18] 2.4% 2.2% 0.13 [0.02, 0.95] 0.25 [0.03, 2.13] 0.34 [0.05, 2.45] Liang J 2018 40 84 40 36 36 88 Wang Y 2017 Subtotal (95% CI) 18.6% 88 Total events 11 23 Heterogeneity: Tau\* = 2.42; Chi\* = 10.18, df = 2 (P = 0.006); I\* = 80% Test for overall effect: Z = 1.08 (P = 0.28) 399 100.0% Total (95% CI) 402 0.62 [0.44, 0.88] Total events 139 204 Heterogeneity: Tau<sup>2</sup> = 0.18; Chi<sup>2</sup> = 41.07, df = 9 (P < 0.00001); I<sup>2</sup> = 78% 0.01 10 100 0.1 Test for overall effect: Z = 2.72 (P = 0.007) Test for subgroup differences: Chi<sup>2</sup> = 0.39, df = 1 (P = 0.53), P = 0% Favours [Kanglaite] Favours [Chemotherapy] Kanglaite Chemotherapy **Risk Ratio Risk Ratio** В Study or Subgroup Events Total Events Total Weight M-H, Random, 95% CI M-H, Random, 95% CI 2.3.1 GP Chen Y 2018 15 18 51 11.0% 0.83 [0.47, 1.47] 0.82 [0.57, 1.18] 0.62 [0.41, 0.93] Guan XQ 2009 9 12 11 12 13.4% 35 35 35 35 12.9% Huang ZB 2010 18 26 23 Liu JQ 2011 8 0.35 [0.18, 0.67] Wang L 2014 Wang Y 2017 7.8% 0.70 [0.29, 1.67] 0.25 [0.03, 2.13] 7 43 10 43 36 36 49 13 Yan QH 2018 12 49 9.7% 0.92 [0.47, 1.82] Yao J 2017 Zhang MM 2019 15.7% 3.5% 60 70 50 65 67 0.88 [0.80, 0.98] 0.40 [0.08, 1.97] 0.70 [0.53, 0.93] 50 2 5 4 Subtotal (95% CI) 381 378 86.2% Total events 130 175 Heterogeneity: Tau<sup>2</sup> = 0.08; Chi<sup>2</sup> = 21.09, df = 8 (P = 0.007); I<sup>2</sup> = 62% Test for overall effect: Z = 2.47 (P = 0.01) 2.3.2 other Jia JN 2018 10 31 26 8 31 11.4% 0.38 [0.23, 0.66] Liang J 2018 Subtotal (95% CI) 40 40 2.4% 0.13 [0.02, 0.95] 0.32 [0.14, 0.75] 1 71 11 34 Total events Heterogeneity: Tau\* = 0.14; Chi\* = 1.25, df = 1 (P = 0.26); I\* = 20% Test for overall effect: Z = 2.63 (P = 0.009) Total (95% CI) 452 449 100.0% 0.61 [0.44, 0.86] Total events 141 209 Heterogeneity: Tau<sup>2</sup> = 0.19; Chi<sup>2</sup> = 43.25, df = 10 (P < 0.00001); l<sup>2</sup> = 77% 0.01 0.1 10 100 Test for overall effect: Z= 2,84 (P = 0.004) Test for subgroup differences:  $Chi^{\mu}$ = 2.94. df = 1 (P = 0.09). P = 66.0% Favours [Kanglaite] Favours [Chemotherapy] Kanglaite Chemotherapy С **Risk Ratio Risk Ratio** Study or Subgroup Events Total Events Total Weight M-H, Random, 95% CI M-H, Random, 95% CI 2.4.1 2-cycle Guan XQ 2009 9 13.4% 0.82 [0.57, 1.18] 12 11 12 Huang ZB 2010 Jia JN 2018 35 31 12.9% 11.4% 0.62 [0.41, 0.93] 0.38 [0.23, 0.66] 16 10 35 26 26 31 Liano J 2018 1 40 8 40 2.4% 0.13 [0.02, 0.95] Liu JQ 2011 35 43 8 23 35 10.0% 0.35 (0.18, 0.67) Wang L 2014 10 43 7.8% 0.70 (0.29, 1.67) Yao J 2017 60 70 65 67 15.7% 0.88 (0.80, 0.98) Subtotal (95% CI) 266 263 73.6% 0.56 [0.35, 0.90] 111 169 Total events Heterogeneity: Tau<sup>2</sup> = 0.29; Chi<sup>2</sup> = 46.54, df = 6 (P < 0.00001); P = 87% Test for overall effect: Z = 2.40 (P = 0.02) 2.4.2 other 0.83 [0.47, 1.47] Chen Y 2018 15 11.0% 18 51 51 2.2% 9.7% 0.25 [0.03, 2.13] 0.92 [0.47, 1.82] Wang Y 2017 1 36 4 36 Yan QH 2018 12 49 13 49 Zhang MM 2019 3.5% 2 50 5 50 0.40 [0.08, 1.97] 0.79 [0.52, 1.19] Subtotal (95% CI) 186 186 26.4% Total events 30 40 Heterogeneity: Tau\* = 0.00; Chi\* = 2.10, df = 3 (P = 0.55); I\* = 0% Test for overall effect: Z = 1.13 (P = 0.26) Total (95% CI) 452 0.61 [0.44, 0.86] 449 100.0% Total (95% CI) 402 446 1000.5 Total events 141 209 Heterogeneity: Tau<sup>2</sup> = 0.19; Chi<sup>2</sup> = 43.25; df = 10 (P < 0.00001); I<sup>2</sup> = 77% 0.01 10 0.1 100 Test for overall effect: Z = 2.84 (P = 0.004) Test for subgroup differences: Chi<sup>#</sup> = 1.10. df = 1 (P = 0.29). P = 9.3%Favours [Kanglaite] Favours [Chemotherapy] Kanglaite Chemotherapy **Risk Ratio Risk Ratio** D Study or Subgroup **Events Total Events** Total Weight M-H. Random, 95% CI M-H, Random, 95% CI 2.7.1 yes Guan XQ 2009 g 11 12 13.4% 0.82 (0.57, 1.18) 12 35 35 26 23 35 35 12.9% 0.62 [0.41, 0.93] 0.35 [0.18, 0.67] Huang ZB 2010 16 Liu JQ 2011 87 Wang L 2014 Yao J 2017 Subtotal (95% CI) 0.70 [0.29, 1.67] 0.86 [0.80, 0.98] 0.67 [0.46, 0.99] 43 10 43 7.8% 60 70 65 15.7% 195 192 59.8% Total events 100 135 Heterogeneity: Tau<sup>2</sup> = 0.14; Chi<sup>2</sup> = 19.93, df = 4 (P = 0.0005); l<sup>2</sup> = 80% Test for overall effect: Z = 1.99 (P = 0.05) 2.7.2 unclear 0.83 [0.47, 1.47] 0.38 [0.23, 0.66] Chen Y 2018 51 31 11.0% 15 51 18 11.4% Jia JN 2018 10 31 26 Liang J 2018 40 1 8 40 0.13 [0.02, 0.95] Wang Y 2017 Yan QH 2018 36 49 2.2% 0.25 [0.03, 2.13] 0.92 [0.47, 1.82] 36 49 12 13 Zhang MM 2019 Subtotal (95% CI) 2 50 5 50 3.5% 0.40 (0.08, 1.97) 257 257 40.2% 0.55 [0.33, 0.90] 41 74 Total events Heterogeneity: Tau\* = 0.14; Chi\* = 8.74, df = 5 (P = 0.12); I\* = 43% Test for overall effect: Z = 2,38 (P = 0.02)

|   | Total (95% CI)                    |              | 452       |               | 449       | 100.0%    | 0.61 [0.44, 0.86]   |       | +                    |   |
|---|-----------------------------------|--------------|-----------|---------------|-----------|-----------|---------------------|-------|----------------------|---|
|   | Total events                      | 141          |           | 209           |           |           |                     |       |                      |   |
|   | Heterogeneity: Tau <sup>2</sup> = | = 0.19; Ch   | = 43.     | 25, df = 10   | (P < 0.0  | 0001); P: | = 77%               | 10.04 | di.                  | 10 100  |
|   | Test for overall effect:          | Z= 2.84      | P = 0.0   | 04)           |           |           |                     | 0.01  | U.1                  | Tu 100  |
|   | Test for subaroup dif             | ferences:    | Chi#=     | 0.41. df = 1  | (P = 0.5) | 2). F= 0  | 6                   |       | Payouts (Marigiaite) | Favours [Chemotherapy]  |
| E |                                   | Kangk        | tite      | Chemoth       | erapy     |           | Risk Ratio          |       | Risk                 | Ratio   |
| - | Study or Subgroup                 | Events Total |           | Events        | Total     | Weight    | M-H, Random, 95% CI | -     | M-H, Rand            | om, 95% CI  |
|   | 2.6.1 YES                         |              |           |               |           |           |                     |       |                      | the second se |
|   | Guan XQ 2009                      | 9            | 12        | 11            | 12        | 13.4%     | 0.82 [0.57, 1.18]   |       |                      | -   |
|   | Jia JN 2018                       | 10           | 31        | 26            | 31        | 11.4%     | 0 38 [0.23, 0.66]   |       |                      |   |
|   | Liang J 2018                      | 1            | 40        | 8             | 40        | 2.4%      | 0.13 (0.02, 0.95)   | -     |                      | C   |
|   | Liu JQ 2011                       | 8            | 35        | 23            | 35        | 10.0%     | 0.35 [0.18, 0.67]   |       |                      |   |
|   | Wang L 2014                       | 7            | 43        | 10            | 43        | 7.8%      | 0.70 [0.29, 1.67]   |       |                      |   |
|   | Yan QH 2018                       | 12           | 49        | 13            | 49        | 9.7%      | 0.92 [0.47, 1.82]   |       |                      |   |
|   | Subtotal (95% CI)                 |              | 210       |               | 210       | 54.7%     | 0.55 [0.35, 0.87]   |       | •                    |   |
|   | Total events                      | 47           |           | 91            |           |           |                     |       |                      |   |
|   | Heterogeneity: Tau <sup>2</sup> = | = 0.18; Ch   | = 13.     | 73, df = 5 (l | P=0.02    | ;F= 649   | 6                   |       |                      |   |
|   | Test for overall effect           | Z= 2.56      | (P = 0.0) | )1)           |           |           |                     |       |                      |   |
|   | 2.6.2 Unclear                     |              |           |               |           |           |                     |       |                      |   |
|   | Chen Y 2018                       | 15           | 51        | 18            | 51        | 11.0%     | 0.83 (0.47, 1.47)   |       | 1                    | -   |
|   | Huang ZB 2010                     | 16           | 35        | 26            | 35        | 12.9%     | 0.62 [0.41, 0.93]   |       |                      |   |
|   | Wang Y 2017                       | 1            | 36        | 4             | 36        | 2.2%      | 0.25 [0.03, 2.13]   |       |                      |   |
|   | Yao J 2017                        | 60           | 70        | 65            | 67        | 15.7%     | 0.88 [0.80, 0.98]   |       |                      |   |
|   | Zhang MM 2019                     | 2            | 50        | 5             | 50        | 3.5%      | 0.40 [0.08, 1.97]   |       |                      |   |
|   | Subtotal (95% CI)                 |              | 242       |               | 239       | 45.3%     | 0.74 [0.52, 1.05]   |       | •                    |   |
|   | Total events                      | 94           |           | 118           |           |           |                     |       |                      |   |
|   | Heterogeneity: Tau* =             | = 0.07; Ch   | "= 8.9    | 4, df = 4 (P  | = 0.06);  | 1= 55%    |                     |       |                      |   |
|   | Test for overall effect           | Z=1.70       | (P = 0,0  | )9)           |           |           |                     |       |                      |   |
|   | Total (95% CI)                    |              | 452       |               | 449       | 100.0%    | 0.61 [0,44, 0.86]   |       | +                    | 1   |
|   | Total events                      | 141          |           | 209           |           |           |                     |       |                      |   |
|   | Heterogeneity: Tau <sup>2</sup> = | = 0.19; Ch   | = 43.     | 25, df = 10   | (P < 0.0  | 0001); F= | = 77%               | 10.04 | di                   | 10 100  |
|   | Test for overall effect:          | Z= 2.84      | P=0.0     | 04)           |           |           |                     | 0.01  | U.1                  | TU 100  |
|   | Test for subgroup dif             | ferences:    | Chi= I    | 0.97. df = 1  | (P = 0.3) | 2). F= 0  | 8                   |       | Pavours (Kanglaite)  | Payours (Chemotherapy)  |

Figure S4 Subgroup analysis of leukopenia for each variable. (A) Subgroup analysis of leukopenia for Kanglaite dose; (B) subgroup analysis of leukopenia for type of first-line platinum-based chemotherapy; (C) subgroup analysis of leukopenia for cycle of chemotherapy; (D) subgroup analysis of leukopenia for evaluation criteria; (E) subgroup analysis of leukopenia for supportive treatment.

| Question: Should Kanglaite combined with chemotherapy vs chemotherapy be used for NSCLC? |                      |  |   |                                       |              |  |                               |   |                           |                              |  |   |                  |
|--|----------------------|--|---|---------------------------------------|--------------|--|-------------------------------|---|---------------------------|------------------------------|--|---|------------------|
| Quality assessment   |                      |  |   |                                       |              |  | Summary of Findings           |   |                           |                              |  |   |                  |
| Participants   | Risk of              | Inconsistency  | Indirectness  | Imprecision                           | Publication  | Overall quality of                           | Study ev                      | vent rates (%)                                  | Relative effect           | Anticipated absolute effects |  |   |                  |
| (studies)<br>Follow up   | bias                 |  |   |                                       | bias         | evidence                                     | With<br>Chemotherapy          | With Kanglaite<br>combined with<br>chemotherapy | (95% CI)                  | Risk with Chemotherapy       | Risk difference with Kanglaite<br>combined with chemotherapy<br>(95% Cl) |   |                  |
|  |                      |  |   |                                       | CR+PF        | R (CRITICAL OUTCOM                           | 1E; assessed with: follow up) |   |                           |                              |  |   |                  |
| 2577   | very                 | very no serious<br>serious <sup>1,2</sup> inconsistency    | no serious no serious<br>nconsistency indirectness      | no serious                            | undetected   | ⊕⊕⊝⊝<br>L O)4/12                             | 415/1286                      | 588/1291<br>(45.5%)                             | RR 1.41<br>(1.28 to 1.56) | Stud                         | ly population  |   |                  |
| 1 to 4 weeks   | serious"-            |  |   | imprecision                           |              | due to risk of bias                          | (32.3%)                       |   |                           | 323 ORR per 1000             | 132 more ORR per 1000<br>(from 90 more to 181 more)                      |   |                  |
|  |                      |  |   |                                       |              |  |                               |   |                           | Ν                            | Moderate   |   |                  |
|  |                      |  |   |                                       |              |  |                               |   |                           | 333 ORR per 1000             | 137 more ORR per 1000<br>(from 93 more to 186 more)                      |   |                  |
|  |                      |  |   |                                       | CR+PR+       | SD (CRITICAL OUTCO                           | OME; assessed wit             | th: follow up)                                  |                           |                              |  |   |                  |
| 2493   | very                 | y no serious<br>us <sup>1,2</sup> inconsistency            | no serious  | no serious                            | undetected   | ⊕⊕⊝⊝   | 856/1244                      | 1000/1249<br>(80.1%)                            | RR 1.16<br>(1.11 to 1.22) | Study population             |  |   |                  |
| (31 studies)<br>1 to 4 weeks   | serious              |  | / indirectness  | imprecision                           |              | LOW <sup>1,2</sup><br>due to risk of bias    | (68.8%)                       |   |                           | 688 DCR per 1000             | 110 more DCR per 1000<br>(from 76 more to 151 more)                      |   |                  |
|  |                      |  |   |                                       |              |  |                               |   |                           | Moderate                     |  |   |                  |
|  |                      |  |   |                                       |              |  |                               |   |                           | 673 DCR per 1000             | 108 more DCR per 1000<br>(from 74 more to 148 more)                      |   |                  |
|  |                      |  |   |                                       | KPS (IN      | IPORTANT OUTCOM                              | E; assessed with:             | KPS scale)                                      |                           |                              |  |   |                  |
| 737  | serious <sup>1</sup> | no serious no serio<br>inconsistency indirectr             | no serious no seri<br>cy indirectness impreci           | no serious                            | undetected   | ed 000000000000000000000000000000000000      | 102/368<br>(27.7%)            | 186/369<br>(50.4%)                              | RR 1.82<br>(1.51 to 2.19) | Study population             |  |   |                  |
| (10 studies)<br>1 to 4 weeks   |                      |  |   | imprecision                           |              |  |                               |   |                           | 277 KPS per 1000             | 227 more KPS per 1000<br>(from 141 more to 330 more)                     |   |                  |
|  |                      |  |   |                                       |              |  |                               |   |                           | Moderate                     |  |   |                  |
|  |                      |  |   |                                       |              |  |                               |   |                           |                              |  |   | 673 KPS per 1000 |
|  |                      |  |   |                                       | nausea and v | omiting (CRITICAL OL                         | JTCOME; assesse               | d with: follow up)                              |                           |                              |  |   |                  |
| 672  | serious1             | no serious   | no serious  | serious <sup>3</sup>                  | undetected   | ⊕⊕⊝⊝   | 168/333                       | 111/339   | RR 0.58                   | Study population             |  |   |                  |
| (9 studies)<br>1 to 4 weeks  |                      | inconsistency  | indirectness  |                                       |              | due to risk of bias,<br>imprecision          | (50.5%)                       | (32.7%)   | (32.7%)                   | (0.42 to 0.81)               | 505 NV per 1000  | 212 fewer NV per 1000<br>(from 96 fewer to 293 fewer) |                  |
|  |                      |  |   |                                       |              |  |                               |   | Low                       |                              |  |   |                  |
|  |                      |  |   |                                       |              |  |                               |   |                           | 400 NV per 1000              | 168 fewer NV per 1000<br>(from 76 fewer to 232 fewer)                    |   |                  |
| leukopenia (CRITICAL OUTCOME; assessed with: follow up)                                  |                      |  |   |                                       |              |  |                               |   |                           |                              |  |   |                  |
| 901  | serious <sup>1</sup> | erious <sup>1</sup> no serious no s<br>inconsistency indir | no serious no serious no<br>consistency indirectness im | ious no serious u<br>ness imprecision | undetected   | @@@@   | 209/449<br>(46.5%)            | 141/452<br>(31.2%)                              | RR 0.61<br>(0.44 to 0.86) | Stud                         | ly population  |   |                  |
| (10 studies)<br>1 to 4 weeks   |                      |  |   |                                       |              | MODERATE <sup>1</sup><br>due to risk of bias |                               |   |                           | 465 LP per 1000              | 182 fewer LP per 1000<br>(from 65 fewer to 261 fewer)                    |   |                  |
|  |                      |  |   |                                       |              |  |                               |   |                           | Moderate                     |  |   |                  |
|  |                      |  |   |                                       |              |  |                               |   |                           | 673 LP per 1000              | 262 fewer LP per 1000<br>(from 94 fewer to 377 fewer)                    |   |                  |

<sup>1</sup> lack of blinding; <sup>2</sup> selective bias; <sup>3</sup> regards nausea and vomit as two different symptoms to rate

Figure S5 Summary of evidence.