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Protocol

3 **Main inclusion criteria:**

4 Age ≥ 18 , Pathologically identified non-small cell lung cancer

5 Adenocarcinoma, No-drive gene mutations (with EGFR, ALK, ROS1 included detected by next
6 generation sequencing, NGS)

7 Squamous cell carcinoma, there is no need to perform gene detection but will be retrospectively
8 detected by NGS

9 Progression disease after second line treatment

10 Patients have not been administrated with Vinorelbine or apatinib

11 PS score ≤ 2

12 Estimated survival time more than 3 months

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14 **Main exclusion criteria:**

15 Patients received treatment of apatinib or Vinorelbine before

16 EGFR, ALK or ROS1 mutation positive for Lung Adenocarcinoma patients

17 EGFR, ALK or ROS1 mutation positive for Squamous cell carcinoma if the patients have
18 performed gene detection by NGS

19 Patients with contraindication of chemotherapy

20 Pregnant or breast feeding women

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22 **Outcome Measures:**

23 Primary Outcome: Objective Response Rate (ORR)

24 ORR means the best response status during the whole treatment line.

25 Secondary Outcome Measures: Progression free survival (PFS), OS (Overall survival) and safety

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27 **Sample size calculation:**

28 We have chosen a fixed sample size of 30

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30 **Statistical analysis:**

31 SPSS (version 22.0) was used for statistical analyses. Proportion of responders were calculated

32 with the 95% confidence intervals (CI) using the Clopper-Pearson method. Kaplan-Meier method

33 was used to estimate the median duration of response and PFS with corresponding 95% CIs.