**Effect of Huaier Granule on Recurrence after Curative Resection of HCC:**

**A Multicenter, Randomized Clinical Trial**

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**Supplemental Material**

Site distribution and patient number were as follows: Tongji Hospital of HUST (69), Wuhan; Southwest University Hospital (25), Chongqing; the First Affiliated Hospital of Harbin Medical University (23), Harbin; the Third Affiliated Hospital of Harbin Medical University (44), Harbin; Shengli Hospital Affiliated to Anhui Medical University (36), Hefei; Taihe Hospital Affiliated to Hubei University of Medicine (20), Shiyan; Wuhan Union Hospital of HUST (39), Wuhan; Cancer Hospital Affiliated to Xinjiang Medical University (27), Urumqi; the First Affiliated Hospital of Shihezi University School of Medicine (26), Shihezi; Cancer Hospital of Chinese Academy of Medical Sciences (51), Beijing; Cancer Hospital Affiliated to Guangxi Medical University (45), Nanning; the First Affiliated Hospital of Suzhou Medical University (19), Suzhou; Guangxi Liuzhou Worker’s Hospital (22), Liuzhou; the First Affiliated Hospital of Sun Yat-Sen University (77), Guangzhou; Chinese Arsenal and Police (Wujing) General Hospital (6), Beijing; the First Affiliated Hospital of Xinjiang Medical University (27), Urumqi; the First People’s Hospital (53), Foshan; the Second People’s Hospital (4), Shenzhen; Tanjin No.3 Hospital (17), Tianjin; Beijing Ditan Hospital Affiliated to Capital Medical University (23), Beijing; Hubei Cancer Hospital (2), Wuhan; China and Japan Union Hospital of Jilin Hospital (25), Changchun; Shengjing Hospital of China Medial University (25), Shenyang; Chinese PLA General Hospital of Medical School of Chinese PLA (32), Beijing; West China Hospital of Sichuan University (36), Chengdu; Nanjing Drum Tower Hospital Affiliated to Nanjing University Medical School (45), Nanjing; the Second Xiangya Hospital of Central South University (9), Changsha; Shanghai Changzheng Hospital (23), Shanghai; Beijing Cancer Hospital (24), Beijing; Henan Cancer Hospital Affiliated to Zhengzhou University (22), Zhengzhou; the Second Affiliated Hospital of Zhejiang University School of Medicine (4), Hangzhou; Hospital of Infectious and Contagious Diseases Affiliated to Fujian Medical University (16), Fuzhou; Shanghai Eastern Hepatobiliary Surgery Hospital (34), Shanghai; Tianjin Medical University Cancer Institute & Hospital (TMUCIH) (13), Tianjin; the Affiliated Hospital of Guizhou Medical University (6), Guiyang; Fujian Cancer Hospital (9), Fuzhou; Changzhou TCM Hospital (1), Changzhou; Lingzhe Cancer Hospital (7), Lingzhe; Third People’s Hospital of Changzhou (16), Changzhou.

**Supplementary Table 1.** Schoenfeld residual test for assessment of proportional hazard assumption

|  |  |  |  |
| --- | --- | --- | --- |
|  | RFS | OS | ERFS |
| Correlation coefficient | -0.006 | -0.056 | 0.058 |
| *P*-value | 0.9002 | 0.6732 | 0.4981 |

**Supplementary Table 2.** Baseline disease characteristics between Huaier and control groups

|  |  |  |  |
| --- | --- | --- | --- |
|  | Huaier group (N=686) | Control group  (N=316) | *P*-value |
| BMI | 23.78±3.56 | 23.44±3.30 | 0.1901 |
| ALT (U/L) | 45.82±58.59 | 49.30±80.11 | 0.4941 |
| AST (U/L) | 41.21±45.87 | 44.83±55.85 | 0.3212 |
| Total bilirubin (umol/L) | 16.29±25.02 | 14.33±6.40 | 0.0565 |
| Haemoglobin (g/L) | 142.10±22.15 | 139.90±18.10 | 0.1030 |
| Blood platelet count (109/L) | 161.90±69.91 | 152.20±66.38 | 0.0393 |
| Absolute neutrophil count (109/L) | 3.90±2.41 | 3.72±2.13 | 0.2224 |
| Serum creatinine (umol/L) | 74.00±22.91 | 74.52±21.68 | 0.7380 |

The levels of BMI, ALT, AST, total bilirubin, haemoglobin, platelet count, neutrophil count, and serum creatinine were compared between Huaier and control groups at the time of randomization. Data represent the means ± SD. P values were calculated using unpaired Student's t test. *P* < 0.05 was considered as a statistical significance.

**Supplementary Table 3.** CMH-*X*2 test for comparison of local recurrence between Huaier and control groups.

|  |  |  |
| --- | --- | --- |
|  | Huaier（N=686） | Control（N=316） |
| Recurrence-free, n(%) | 472 (68.80) | 191 (60.44) |
| Intrahepatic recurrence\*, n(%) | 214 (31.20) | 125 (39.56) |
| CMH-χ2 | 6.7502 | |
| *P* | .0094 | |
| Difference, 95% CI | 8.36 (1.95,14.77) | |

\*: Among those with local recurrence, 15 patients in Huaier group and 7 patients in control group had both intrahepatic and extrahepatic recurrence, respectively.

**Supplementary Table 4.** Breslow-Day test for the homogeneity across centers on recurrence, death, and extrahepatic recurrence

