

Supplementary Tables

SUPPL. TABLE I- Cell material used for the study

| Type of cell line | ATCC generic name | Disease | Disease markers |
|-------------------|-------------------|--|-----------------|
| 3T3 | ATCC1®CRL-1658™ | Normal Rat muscle | N/A |
| HT-29 | ATCC1® HTB-38™ | Colorectal adenocarcinoma | N/A |
| HCT-15 | ATCC® CCL-225™ | Colorectal adenocarcinoma, dukes type C/ resistant variant | P-gp |

N/A: Not Applicable, P-gp: P-glycoprotein

SUPPL.TABLE II- Analysis for combination index range

| Range of combination index | Description |
|----------------------------|------------------------|
| <0.1 | Very strong synergism |
| 0.1-0.3 | Strong synergism |
| 0.3-0.7 | Synergism |
| 0.7-0.85 | Moderate synergism |
| 0.85-0.90 | Slight synergism |
| 0.90-1.10 | Nearly additive |
| 1.10-1.20 | Slight antagonism |
| 1.20-1.45 | Moderate antagonism |
| 1.45-3.3 | Antagonism |
| 3.3-10 | Strong antagonism |
| >10 | Very strong antagonism |

TABLE III- The retention factors of the separated components of the *S. birrea* ethanol extract using different mobile phases

| Mobile phases | Compound | Retention factor |
|--|----------|------------------|
| DCM: Hexane: Ethyl Acetate | 1 | 0.47 |
| | 2 | 0.65 |
| | 3 | 0.93 |
| | 4 | 0.99 |
| | 1 | 0.2 |
| Benzene: methanol: Ammonium hydroxide | 2 | 0.35 |
| | 3 | 0.54 |
| | 4 | 0.88 |
| | 5 | 0.95 |

Suppl. Table IV. The IC₅₀ and combination index values of the DOX and VER on HT29 cells, n=3.

| Drug/Combo | CI value | Dose VER | Dose DOX | DRI VER | DRI DOX |
|------------------|----------|----------|----------|---------|---------|
| IC ₅₀ | | | | | |
| VER | | 3.04 | | 2.3 | |
| DOX | | | 0.45 | | 1.7 |
| DOX + VER | 1.00 | 1.31 | 0.26 | | |
| IC ₇₅ | | | | | 3.5 |

| | | | | | |
|------------------------|------|-------|------|-----|------|
| VER | | 23.06 | | 7.8 | |
| DOX | | | 2.07 | | |
| DOX + VER | 0.41 | 2.94 | 0.59 | | |
| IC₉₀ | | | | | |
| VER | | 175.1 | | 26 | |
| DOX | | | 9.40 | | 7 |
| DOX + VER | 0.17 | 6.61 | 1.32 | | |
| IC₉₅ | | | | | |
| VER | | 695.4 | | 60 | |
| DOX | | | 26.7 | | 11.4 |
| DOX+ VER | 0.10 | 11.5 | 2.29 | | |

Suppl. Table V. The IC and combination index values of the DOX and SBE extract on HT29 cells, n=3.

| Drug/Combination | CI value | Dose-SBE | Dose-DOX | DRI-SBE | DRI-DOX |
|------------------------|----------|----------|----------|---------|---------|
| IC₅₀ | | | | | |
| SBE | | 140.09 | | 0.21 | |
| DOX | | | 0.35 | | 0.20 |
| SBE + DOX | 9.6 | 669.89 | 1.70 | | |
| IC₇₅ | | | | | |
| SBE | | 689.90 | | 0.14 | |
| DOX | | | 1.26 | | 0.10 |
| SBE + DOX | 17.1 | 4969.46 | 12.6 | | |
| IC₉₀ | | | | | |
| SBE | | 3397.65 | | 0.09 | |
| DOX | | | 4.5 | | 0.04 |
| SBE + DOX | 31.6 | 36865.0 | 93.6 | | |
| IC₉₅ | | | | | |
| SBE | | 10048.5 | | 0.07 | |
| DOX | | | 10.7 | | 0.02 |
| SBE + DOX | 48.5 | 144061 | 365.9 | | |

Suppl. Table VI. The IC and combination index values of DOX and SBW extract on HT29 cells, n=3.

| Drug/Combination | CI value | Dose S.B H ₂ O | Dose DOX | DRI SBW | DRI DOX |
|------------------------|----------|---------------------------|----------|---------|---------|
| IC₅₀ | | | | | |
| SBW | | 208.0 | | 0.3 | |
| DOX | | | 0.47 | | 0.4 |
| SBW + DOX | 2.5 | 275.4 | 0.57 | | |
| IC₇₅ | | | | | |
| SBW | | 940.70 | | 0.21 | |
| DOX | | | 2.04 | | 0.50 |
| SBW + DOX | 0.97 | 467.39 | 0.97 | | |
| IC₉₀ | | | | | |
| SBW | | 4253.9 | | 0.13 | |
| DOX | | | 8.89 | | 0.59 |
| SBW + DOX | 0.37 | 793.13 | 1.64 | | |
| IC₉₅ | | | | | |
| SBW | | 11871.9 | | 0.09 | |
| DOX | | | 24.18 | | 0.66 |
| SBW + DOX | 0.19 | 1136.5 | 2.36 | | |

Suppl. Table VII. The IC and combination index values of DOX and VER on HCT15 cells, n=3.

| Drug/Combination | CI value | Dose VER | Dose DOX | DRI VER | DRI DOX |
|------------------------|----------|----------|----------|---------|---------|
| IC₅₀ | | | | | |
| VER | | 6.63 | | 2.64 | |
| DOX | | | 0.38 | | 0.46 |
| DOX + VER | 1.33 | 2.99 | 0.33 | | |
| IC₇₅ | | | | | |
| VER | | 217.40 | | 355.6 | |
| DOX | | | 6.11 | | 1.1 |
| DOX + VER | 0.34 | 14.82 | 1.66 | | |
| IC₉₀ | | | | | |
| VER | | 7127.23 | | 47807.7 | |
| DOX | | | 97.73 | | 3.0 |
| DOX + VER | 0.09 | 73.44 | 8.25 | | |
| IC₉₅ | | | | | |
| VER | | 7652.5 | | 1340243 | |
| DOX | | | 643.6 | | 4.2 |
| DOX+ VER | 0.04 | 218.07 | 24.50 | | |

Suppl. Table VIII. The IC and combination index values of DOX and SBE extract on HCT15 cells.

| Drug/Combination | CI value | Dose SB-EE | Dose DOX | DRI SB- EE | DRI DOX |
|-------------------------|-----------------|-------------------|-----------------|-------------------|----------------|
| IC₅₀ | | | | | |
| SBE | | 49.35 | | 0.54 | |
| DOX | | | 0.68 | | 0.51 |
| SBE + DOX | 3.75 | 89.86 | 1.32 | | |
| IC₇₅ | | | | | |
| SBE | | 256.47 | | 7.37 | |
| DOX | | | 4.12 | | 8.05 |
| SBE + DOX | 0.26 | 34.86 | 0.51 | | |
| IC₉₀ | | | | | |
| SBE | | 1332.76 | | 98.5 | |
| DOX | | | 24.87 | | 125.1 |
| SBE + DOX | 0.01 | 13.53 | 0.20 | | |
| IC₉₅ | | | | | |
| SBE | | 4088.34 | | 578.37 | |
| DOX | | | 84.43 | | 809.18 |
| SBE + DOX | 0.00297 | 7.11 | 0.10 | | |

Suppl. Table IX. The IC and combination index values of DOX and SBW extract on HCT15 cells, n=3.

| Drug/Combination | CI value | Dose SBW | Dose DOX | DRI SBW | DRI DOX |
|-------------------------|-----------------|-----------------|-----------------|----------------|----------------|
| IC₅₀ | | | | | |
| SBW | | 413.41 | | 0.23 | |
| DOX | | | 1.54 | | 0.23 |
| SBW + DOX | 5.30 | 1272.60 | 3.62 | | |
| IC₇₅ | | | | | |
| SBW | | 1689.40 | | 0.15 | |
| DOX | | | 11.33 | | 0.18 |
| SBW + DOX | 6.66 | 7903.88 | 22.50 | | |
| IC₉₀ | | | | | |
| SBW | | 6615.65 | | 0.07 | |
| DOX | | | 83.28 | | 0.14 |
| SBW + DOX | 10.00 | 49089.5 | 139.74 | | |
| IC₉₅ | | | | | |
| SBW | | 16741.3 | | 0.05 | |
| DOX | | | 323.40 | | 0.11 |
| SBW + DOX | 11.65 | 169998 | 483.91 | | |

Supplementary Figures



(A)



(B)

Suppl. Fig. 1 - TLC Chemical fingerprint of the SBE extract. (A) DCM: EA: HEX (B) Benzene: MeOH: NH₃OH at 366 nm.