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## Supplemental Table 1. Search Strategy

| Deadline N | May 18, 2023   |
|------------|--|
| CINAHL     | (colorectal) OR (colon) OR (rectal) OR (colostomy)   |
| PLUS       | AND  |
|            | (cancer) OR (tumor) OR (neoplasm) OR (carcinoma)   |
|            | AND  |
|            | (cognitive dysfunction) OR (cognitive impairment) OR (cognitive disorder) OR (cognitive    |
|            | decline) OR (cognitive complaint)  |
| Cochrane   | #1 (colorectal):ti,ab,kw OR (colon):ti,ab,kw OR (rectal):ti,ab,kw OR (colostomy):ti,ab,kw  |
| Library    | #2 (cancer):ti,ab,kw OR (tumor):ti,ab,kw OR (neoplasm):ti,ab,kw OR (carcinoma):ti,ab,kw    |
|            | #3 (cognitive dysfunction):ti,ab,kw OR (cognitive impairment):ti,ab,kw OR (cognitive       |
|            | disorder):ti,ab,kw OR (cognitive decline):ti,ab,kw OR (cognitive complaint):ti,ab,kw       |
|            | #1 AND #2 AND #3   |
| Embase     | #1 'colorectal' OR 'colon' OR 'rectal' OR 'colostomy'                                      |
|            | #2 'cancer' OR 'tumor' OR 'neoplasm' OR 'carcinoma'  |
|            | #3 'cognitive dysfunction' OR 'cognitive impairment' OR 'cognitive disorder' OR 'cognitive |
|            | decline' OR 'cognitive complaint'  |
|            | #1 AND #2 AND #3   |
| PsycINFO   | (colorectal) OR (colon) OR (rectal) OR (colostomy)   |
|            | AND  |
|            |  |

|        | (cancer) OR (tumor) OR (neoplasm) OR (carcinoma)  |
|--------|---|
|        | AND   |
|        | (cognitive dysfunction) OR (cognitive impairment) OR (cognitive disorder) OR (cognitive       |
|        | decline) OR (cognitive complaint)   |
| PubMed | Search terms: (((((colorectal) OR (colon)) OR (rectal)) OR (colostomy)) AND ((((cancer) OR    |
|        | (tumor)) OR (neoplasm)) OR (carcinoma))) AND ((((cognitive dysfunction) OR (cognitive         |
|        | impairment)) OR (cognitive disorder)) OR (cognitive decline)) OR (cognitive complaint))       |
| CNKI   | 'colorectal' + 'colon' + 'rectal' + 'colostomy'   |
|        | AND   |
|        | 'cancer' + 'tumor' + 'neoplasm' + 'carcinoma'   |
|        | AND   |
|        | 'cognitive dysfunction' + 'cognitive impairment' + 'cognitive disorder' + 'cognitive decline' |
|        | + 'cognitive complaint'   |

## Supplemental Table 1. Study Characteristics

| Author  | Study      | Target          | Treatment    | Time       | Outcome    | Main Findings  |
|---------|------------|-----------------|--------------|------------|------------|----------------|
| (year)  | design     | population      | (before      | Points of  | Measures   | and Conclusion |
| Country |            | (Cancer         | and/or       | Assessment | for        | (prevalence of |
|         |            | stage, age,     | after        |            | Cognitio   | cognitive      |
|         |            | education)      | treatment)   |            | n          | impairment,    |
|         |            |                 |              |            | (Objecti   | factors        |
|         |            |                 |              |            | ve and     | contributing   |
|         |            |                 |              |            | subjecti   | to cognitive   |
|         |            |                 |              |            | ve scale)  | impairment,    |
|         |            |                 |              |            |            | interventions  |
|         |            |                 |              |            |            | on cognitive   |
|         |            |                 |              |            |            | impairment)    |
| Andreis | Longitudin | Stage III colon | Undergoing   | T0: before | Objective  | OCI/SCI: not   |
| et al.  | al study   | cancer patients | adjuvant     | any        | scale:     | report         |
|         |            | (n=57)          | chemotherapy |            | MMSE, CDT, |                |

| (2013) | Mean age 58.68 | pharmacologi | Rey Figure  | - No cognitive    |
|--------|----------------|--------------|-------------|-------------------|
| Italy  | (9.62)         | cal agents   | Complex     | impairment in     |
| reary  | Mean education | T1: at the   | (copy and   | time              |
|        | years 9.43     | end of       | recall),    | considered.       |
|        | (3.91)         | adjuvant     | Rey         | -Some significant |
|        |                | treatment    | Auditory    | variation in      |
|        |                | T2: 6 months | Verbal      | tests             |
|        |                | after T1     | Learning    | evaluating        |
|        |                |              | Test, TMT A | verbal memory,    |
|        |                |              | and B       | as Rey Auditory   |
|        |                |              | Subjective  | Verbal Learning   |
|        |                |              | scale: not  | Test.             |
|        |                |              | report      | -Significant      |
|        |                |              |             | variations in     |
|        |                |              |             | information       |
|        |                |              |             | processing        |
|        |                |              |             | speed             |
|        |                |              |             | performances,     |
|        |                |              |             | as TMT A.         |

|          |           |                  |              |              |            | -Adjuvant FOLFOX4  |
|----------|-----------|------------------|--------------|--------------|------------|--------------------|
|          |           |                  |              |              |            | had no effect on   |
|          |           |                  |              |              |            | MMSE test          |
|          |           |                  |              |              |            | performance.       |
|          |           |                  |              |              |            | - FOLFOX4 regimen  |
|          |           |                  |              |              |            | was not able to    |
|          |           |                  |              |              |            | determine          |
|          |           |                  |              |              |            | persistent         |
|          |           |                  |              |              |            | cognitive          |
|          |           |                  |              |              |            | impairment.        |
| CHU Jin- | Cross-    | CRC (n=94)       | Receiving    | Before and   | Objective  | OCI: not report    |
| gai      | sectional | Mean age         | chemotherapy | after        | scale: not | SCI: 62(65.9%)     |
| (2010)   | study     | 48.73(9.2)       |              | chemotherapy | report     | were of cognitive  |
| (2019)   |           | Median education |              |              | Subjective | impairment before  |
| China    |           | years 14         |              |              | scale:     | chemotherapy.      |
|          |           |                  |              |              | FACT-Cog   | After chemotherapy |
|          |           |                  |              |              |            | were 85(90.43%).   |
|          |           |                  |              |              |            | The incidence of   |

|  |  |  | chemotherapy was   |
|--|--|--|--------------------|
|  |  |  | up to 24.47%.      |
|  |  |  | - Chemotherapy     |
|  |  |  | would enhance      |
|  |  |  | the incidence of   |
|  |  |  | patients'          |
|  |  |  | cognitive          |
|  |  |  | impairment, and    |
|  |  |  | between            |
|  |  |  | cognitive          |
|  |  |  | impairment         |
|  |  |  | onset and self-    |
|  |  |  | efficacy level     |
|  |  |  | there was          |
|  |  |  | relativity.        |
|  |  |  | - The incidence of |
|  |  |  | cognitive          |
|  |  |  | impairment in      |
|  |  |  | CRC patients was   |

|          |            |                   |              |               |            | negatively         |
|----------|------------|-------------------|--------------|---------------|------------|--------------------|
|          |            |                   |              |               |            | correlated with    |
|          |            |                   |              |               |            | the level of       |
|          |            |                   |              |               |            | self-efficacy.     |
| Couwenbe | Longitudin | Stage I to IV CRC | Neoadjuvant  | Before        | Subjective | OCI: not report    |
| rg et    | al study   | (n=324)           | chemoradiati | treatment     | scale:     | SCI: at 24 months, |
| al.      |            | Responders        | on or short- | and after 3,  | EORTC QLQ- | 20% of the         |
| al.      |            | (n=272)           | course       | 6, 12, 18,    | C30        | patients reported  |
| (2018)   |            | Median age 65     | radiotherapy | and 24 months |            | worsened cognitive |
| Netherla |            | No-responders     | and          |               |            | functioning        |
| nd       |            | (n=52)            | underwent    |               |            | compared with      |
|          |            | Median age 67     | rectal       |               |            | baseline.          |
|          |            | Education level:  | surgery      |               |            | -Global health and |
|          |            | not report        |              |               |            | cognitive          |
|          |            |                   |              |               |            | functioning        |
|          |            |                   |              |               |            | declined to a      |
|          |            |                   |              |               |            | lesser extent,     |
|          |            |                   |              |               |            | and emotional      |
|          |            |                   |              |               |            | functioning        |

|  |  |  | gradually   |       |
|--|--|--|-------------|-------|
|  |  |  | improved    | over  |
|  |  |  | the time.   |       |
|  |  |  | - Cognitive |       |
|  |  |  | functioning | g and |
|  |  |  | symptoms    | of    |
|  |  |  | fatigue     | and   |
|  |  |  | insomnia    |       |
|  |  |  | remained    |       |
|  |  |  | significant | ly    |
|  |  |  | worse       | in    |
|  |  |  | patients    | on    |
|  |  |  | longer-term | 1.    |
|  |  |  | -Global he  | alth, |
|  |  |  | cognitive   |       |
|  |  |  | function,   | and   |
|  |  |  | emotional   |       |
|  |  |  | function    |       |

|         |            |                 |               |              |             | changed during     |
|---------|------------|-----------------|---------------|--------------|-------------|--------------------|
|         |            |                 |               |              |             | treatment.         |
|         |            |                 |               |              |             | - Patients         |
|         |            |                 |               |              |             | reported lower     |
|         |            |                 |               |              |             | physical, role,    |
|         |            |                 |               |              |             | social, and        |
|         |            |                 |               |              |             | cognitive          |
|         |            |                 |               |              |             | function and       |
|         |            |                 |               |              |             | more insomnia      |
|         |            |                 |               |              |             | and fatigue up     |
|         |            |                 |               |              |             | to 2 years after   |
|         |            |                 |               |              |             | the start of       |
|         |            |                 |               |              |             | treatment.         |
| Cruzado | Longitudin | Stage III and   | Treated with  | Prechemother | Objective   | OCI: a total of 28 |
| et al.  | al Study   | high-risk stage | FOLFOX4 for 6 | ару          | scale: NP   | patients (52%)     |
| (2014)  |            | II colon cancer | months        | Post-        | tests       | showed a decline   |
| (2014)  |            | (n=81)          |               | chemotherapy | - Attention | from pre-          |
| Spain   |            | Mean age 66.96  |               | (approximate | TMT A       | chemotherapy to 6- |
|         |            | (9.52)          |               |              |             | month assessments. |

| Mean education  | ly 5.5 months | -Verbal     | SCI: not report   |
|-----------------|---------------|-------------|-------------------|
| years 6.9 (4.1) | later)        | memory:     | -Adjuvant FOLFOX4 |
|                 | 6 months      | Imm-Mem,    | in patients with  |
|                 | after the end | Imm-Mem-    | colon cancer had  |
|                 | of the last   | Q,          | a negative        |
|                 | cycle of      | Delayed-    | effect on verbal  |
|                 | chemotherapy  | Mem,        | memory, which     |
|                 |               | Delayed-    | deterioration     |
|                 |               | Mem-Q       | was mild and      |
|                 |               | - Verbal    | transient.        |
|                 |               | learning    | -A high decline   |
|                 |               | : LMWT,     | incidence and     |
|                 |               | Spanish     | 13% of patients   |
|                 |               | adaptati    | showed            |
|                 |               | on          | impairment at     |
|                 |               | Barcelon    | the end of        |
|                 |               | a Test      | treatment in      |
|                 |               | - Executive | verbal memory.    |
|                 |               | function    |                   |

|          |        |                  |             |              | :          | -It was not        |
|----------|--------|------------------|-------------|--------------|------------|--------------------|
|          |        |                  |             |              | DSymbol,   | possible to        |
|          |        |                  |             |              | SCWT,      | determine which    |
|          |        |                  |             |              | TMT B      | variables could    |
|          |        |                  |             |              | Subjective | be associated      |
|          |        |                  |             |              | scale: not | with patients'     |
|          |        |                  |             |              | report     | cognitive          |
|          |        |                  |             |              |            | dysfunction.       |
| Dhillon  | Cohort | Stage I to IV    | Adjuvant or | At baseline  | Objective  | OCI: not report    |
|          | study  | CRC (n=343)      | Neoadjuvant | (pre-        | scale: NP  | Subjective         |
| et al.   | Scaay  |                  | Neodajavane | _            |            |                    |
| (2018)   |        | Group 1 (n=289): |             | chemotherapy | tests,     | cognitive          |
|          |        | localized CRC    |             | )            | CANTAB     | impairment: at     |
| Australi |        | Median age 59.0  |             | 6 months     | Subjective | baseline, 18-24%   |
| a        |        | Group 2 (n=73):  |             | 12months     | scale:     | of cancer          |
|          |        | metastatic or    |             | 24 months    | FACT-COG   | survivors reported |
|          |        | recurrent CRC    |             | (group 1)    | version 2  | impairment.        |
|          |        | Median age 55.5  |             |              |            | - No association   |
|          |        | Group 3 (n=72):  |             |              |            | was seen between   |
|          |        | healthy controls |             |              |            | total FACT-COG     |

| Median age 58.5  |  | or PC    | I, and    |
|------------------|--|----------|-----------|
| Education level: |  | neuropsy | ychologi  |
| not report       |  | cal doma | ains.     |
|                  |  | -A weak  | moderate  |
|                  |  | associat | cion was  |
|                  |  | found    | between   |
|                  |  | PCA      | and       |
|                  |  | attentio | on,       |
|                  |  | visual   | memory,   |
|                  |  | or e     | executive |
|                  |  | function | ı.        |
|                  |  | - There  | were      |
|                  |  | signific | cant      |
|                  |  | differer | nces      |
|                  |  | between  | groups    |
|                  |  | in long  | itudinal  |
|                  |  | changes  | on the    |
|                  |  | total    | FACT-COG  |
|                  |  | score,   | and all   |

|  |  |  | subscales from   |
|--|--|--|------------------|
|  |  |  | baseline to 6    |
|  |  |  | months.          |
|  |  |  | -Older           |
|  |  |  | participants     |
|  |  |  | with CRC         |
|  |  |  | reported poorer  |
|  |  |  | cognitive        |
|  |  |  | abilities.       |
|  |  |  | - No association |
|  |  |  | was seen between |
|  |  |  | the FACT-COG     |
|  |  |  | scores and       |
|  |  |  | results of       |
|  |  |  | formal           |
|  |  |  | neuropsychologi  |
|  |  |  | cal testing.     |

| Du et  | Cohort | Stage I     | to IV   | Chemotherapy | At baseline   | Not report | OCI: patients who  |
|--------|--------|-------------|---------|--------------|---------------|------------|--------------------|
| al.    | study  | CRC (n=72,3 | 374)    | and no-      | with up to 17 |            | received           |
| (2013) |        | The         | entire  | chemotherapy | years of      |            | chemotherapy were  |
| (2013) |        | cohort      |         |              | follow-up     |            | 24% significantly  |
| USA    |        | (n=72,374)  |         |              |               |            | more likely to     |
|        |        | Median      | age     |              |               |            | develop drug-      |
|        |        | 73(chemo),  | 78 (no  |              |               |            | induced dementia   |
|        |        | chemo)      |         |              |               |            | compared to those  |
|        |        | The m       | natched |              |               |            | without            |
|        |        | cohort      |         |              |               |            | chemotherapy after |
|        |        | (n=15,921)  |         |              |               |            | adjusting for      |
|        |        | Median age  | 75      |              |               |            | patient and tumor  |
|        |        | Education   | level:  |              |               |            | characteristic.    |
|        |        | not report  |         |              |               |            | SCI: not report    |
|        |        |             |         |              |               |            | - The incidence    |
|        |        |             |         |              |               |            | rate of various    |
|        |        |             |         |              |               |            | cognitive          |
|        |        |             |         |              |               |            | impairments        |
|        |        |             |         |              |               |            | increased with     |

|  |  |  | advanced age  | and  |
|--|--|--|---------------|------|
|  |  |  | higher        |      |
|  |  |  | comorbidity   |      |
|  |  |  | scores but    | was  |
|  |  |  | relatively    |      |
|  |  |  | similar acr   | coss |
|  |  |  | gender and tu | ımor |
|  |  |  | stage.        |      |
|  |  |  | -The status   | of   |
|  |  |  | mood disor    | der  |
|  |  |  | which         |      |
|  |  |  | demonstrated  |      |
|  |  |  | important     |      |
|  |  |  | differences   | in   |
|  |  |  | the           |      |
|  |  |  | associations  |      |
|  |  |  | between       |      |
|  |  |  | chemotherapy  |      |
|  |  |  | use           | and  |

|         |            |                  |              |              |             | cognitive        |
|---------|------------|------------------|--------------|--------------|-------------|------------------|
|         |            |                  |              |              |             | impairments.     |
|         |            |                  |              |              |             | -The receipt of  |
|         |            |                  |              |              |             | chemotherapy     |
|         |            |                  |              |              |             | was associated   |
|         |            |                  |              |              |             | with the         |
|         |            |                  |              |              |             | decreased risk   |
|         |            |                  |              |              |             | of all cognitive |
|         |            |                  |              |              |             | impairments      |
|         |            |                  |              |              |             | combined.        |
| Dwek et | Longitudin | CRC (n=156)      | 50 % of whom | T1: post-    | Objective   | OCI/SCI: not     |
| al.     | al study   | Receive          | received     | surgery but  | scale: MoCA | report           |
|         |            | chemotherapy(n=7 | chemotherapy | prior to     | version 3,  | -A decline in    |
| (2015)  |            | 8)               | , 50 % was   | chemotherapy | NP tests    | cognitive        |
| UK      |            | Non-chemotherapy | nonchemother | treatment    | (HVLT-R,    | functioning can  |
|         |            | surgical         | apy surgical | T2: between  | TMT A and   | be attributed to |
|         |            | patients (n=78)  | patients     | 12 and 14    | B, COWA)    | chemotherapy or  |
|         |            | Age: not report  |              | weeks after  | Subjective  | to disease,      |
|         |            |                  |              | first        | scale:      | surgical or some |

|         |        | Education level: |              | scheduled    | FACT-Cog  | other              |
|---------|--------|------------------|--------------|--------------|-----------|--------------------|
|         |        | not report       |              | chemotherapy | version 3 | confounding        |
|         |        |                  |              | T3: three    |           | factor.            |
|         |        |                  |              | months after |           |                    |
|         |        |                  |              | last         |           |                    |
|         |        |                  |              | scheduled    |           |                    |
|         |        |                  |              | chemotherapy |           |                    |
| Dwek et | Cohort | Stage II and III | Surgery or   | T1: 4 weeks  | Objective | OCI: 45%-55% of    |
| al.     | study  | CRC (n=136)      | chemotherapy | post-surgery | scale: NP | all CRC patients   |
| (2023)  |        | Mean age 61.3    |              | and prior to | tests     | showed cognitive   |
| (2023)  |        | (59.5)           |              | chemotherapy |           | deficits 10 months |
| UK      |        | Median education |              | T2: 12 weeks |           | after surgery.     |
|         |        | years 12         |              | after first  |           | -Memory and        |
|         |        |                  |              | scheduled    |           | executive          |
|         |        |                  |              | chemotherapy |           | function (16% at   |
|         |        |                  |              | T3: 3 months |           | 2SD) were the      |
|         |        |                  |              | after last   |           | cognitive          |
|         |        |                  |              | scheduled    |           | domains most       |
|         |        |                  |              | chemotherapy |           |                    |

|  |  | for patients | frequently        |
|--|--|--------------|-------------------|
|  |  |              |                   |
|  |  | receiving 6  | affected.         |
|  |  | months of    | - Surgery group   |
|  |  | treatment or | showed an         |
|  |  | 6 months     | improvement in    |
|  |  | after last   | test              |
|  |  | scheduled    | performance       |
|  |  | chemotherapy | over time, while  |
|  |  | for patients | the               |
|  |  | receiving 3  | chemotherapy      |
|  |  | months of    | group showed no   |
|  |  | treatment    | improvement.      |
|  |  |              | -The relationship |
|  |  |              | between mood,     |
|  |  |              | fatigue and       |
|  |  |              | cognition was     |
|  |  |              | not strong, with  |
|  |  |              | weak              |
|  |  |              | correlations      |

|  |  |  | found       | between    |
|--|--|--|-------------|------------|
|  |  |  | fatigue     | and        |
|  |  |  | cognitio    | on.        |
|  |  |  | - CRC       | patients   |
|  |  |  | display     |            |
|  |  |  | cognitiv    |            |
|  |  |  | impairme    |            |
|  |  |  | months      |            |
|  |  |  | surgery.    |            |
|  |  |  |             |            |
|  |  |  | - Chemother | rapy did   |
|  |  |  | not         | worsen     |
|  |  |  | cognitiv    | <i>i</i> e |
|  |  |  | impairme    | ent but    |
|  |  |  | did ap      | pear to    |
|  |  |  | slow c      | ognitive   |
|  |  |  | recovery    | 7          |
|  |  |  | relative    | e to       |
|  |  |  | those       |            |

|        |           |                      |              |              |            | undergoing         |
|--------|-----------|----------------------|--------------|--------------|------------|--------------------|
|        |           |                      |              |              |            | surgery only.      |
| Galica | Cross-    | Stage I, II, or      | Undergone    | At baseline  | Objective  | OCI: 25% had       |
| et al. | sectional | III CRC(n=74)        | surgical     | (either      | scale:     | cognitive deficits |
| (2011) | study     | Group A (n = 19):    | resection,   | prior to     | CANTAB     | as determined by   |
| (===/  |           | prechemotherapy      | receive or   | chemotherapy | Subjective | the CANTAB         |
| Canada |           | Group B (n = 18):    | not receive  | administrati | scale: not | results.           |
|        |           | post-surgery         | adjuvant     | on for the   | report     | SCI: not report    |
|        |           | control              | and/or       | cases, or    |            | -No evidence was   |
|        |           | Group C (n = 20):    | neoadjuvant  | within 12    |            | found to suggest   |
|        |           | post-                | chemotherapy | weeks of     |            | that a strong      |
|        |           | chemotherapy         |              | surgery for  |            | relationship       |
|        |           | Group D $(n = 17)$ : |              | the control  |            | exists between     |
|        |           | six-month post-      |              | group)       |            | cognitive          |
|        |           | surgery control      |              | At 6 months  |            | impairment and     |
|        |           | Age: not report      |              | At 12 months |            | psychosocial       |
|        |           | Education level:     |              | At 24 months |            | adjustment, nor    |
|        |           | not report           |              |              |            | that any such      |
|        |           |                      |              |              |            | relationship is    |

|        |     |                  |         |              |           | affected by         |
|--------|-----|------------------|---------|--------------|-----------|---------------------|
|        |     |                  |         |              |           | treatment type      |
|        |     |                  |         |              |           | or time.            |
|        |     |                  |         |              |           | - Cognitive         |
|        |     |                  |         |              |           | changes do not      |
|        |     |                  |         |              |           | influence           |
|        |     |                  |         |              |           | patients'           |
|        |     |                  |         |              |           | relationships       |
|        |     |                  |         |              |           | and functional      |
|        |     |                  |         |              |           | roles, as           |
|        |     |                  |         |              |           | indicated from      |
|        |     |                  |         |              |           | the PAIS-SR.        |
| Gao et | RCT | Stage I and II   | Before  | Before       | Objective | OCI/SCI: not        |
| al.    |     | CRC (n=50)       | surgery | operation    | scale:    | report              |
| (2018) |     | Group C: routine |         | After        | MMSE, TMT | - Compared with the |
|        |     | preoperative     |         | entering the |           | group I, the        |
| China  |     | visit and        |         | operation    |           | MMSE score of       |
|        |     | postoperative    |         | room         |           | the group C was     |
|        |     | follow-up        |         |              |           | significantly       |

| Mean age         | After        | reduced 1, 3 and   |
|------------------|--------------|--------------------|
| 68.6(4.3)        | wakefulness  | 7 days after       |
| Group I:         | During       | surgery.           |
| simultaneous     | follow-up on | - The TMT          |
| psychological    | the 2nd and  | completion time    |
| intervention     | 5th d after  | of the group C     |
| Mean age         | operation    | was                |
| 67.9(3.5)        |              | significantly      |
| Education level: |              | prolonged 1, 3     |
| not report       |              | and 7 days after   |
|                  |              | surgery.           |
|                  |              | - The incidence of |
|                  |              | postoperative      |
|                  |              | cognitive          |
|                  |              | dysfunction in     |
|                  |              | the Group I was    |
|                  |              | significantly      |
|                  |              | decreased 1, 3     |

|        |        |                  |              |              |            | and 7 days after   |
|--------|--------|------------------|--------------|--------------|------------|--------------------|
|        |        |                  |              |              |            | surgery.           |
| Kim et | Cohort | CRC (n=95,303)   | Surgery      | Baseline     | Not report | OCI: the incidence |
| al.    | study  | Mean age         | Chemotherapy | After        |            | rates of cognitive |
| (2021) |        | 64.52(11.9)      | Radiotherapy | chemotherapy |            | impairment were    |
| (2021) |        | Education level: |              |              |            | 22.17 per 1,000    |
| Korea  |        | not report       |              |              |            | person-years in    |
|        |        |                  |              |              |            | chemotherapy non-  |
|        |        |                  |              |              |            | recipients and     |
|        |        |                  |              |              |            | 14.48 per 1,000    |
|        |        |                  |              |              |            | person-years in    |
|        |        |                  |              |              |            | chemotherapy       |
|        |        |                  |              |              |            | recipients.        |
|        |        |                  |              |              |            | SCI: not report    |
|        |        |                  |              |              |            | - The adverse      |
|        |        |                  |              |              |            | effect of          |
|        |        |                  |              |              |            | certain            |
|        |        |                  |              |              |            | chemotherapy       |
|        |        |                  |              |              |            | regimens on        |

|  |  |  | cognition was    |
|--|--|--|------------------|
|  |  |  | more prominent   |
|  |  |  | in older adults. |
|  |  |  | - Neither        |
|  |  |  | chemotherapy     |
|  |  |  | nor              |
|  |  |  | radiotherapy     |
|  |  |  | was positively   |
|  |  |  | associated with  |
|  |  |  | cognitive        |
|  |  |  | impairment.      |
|  |  |  | - The FOLFOX     |
|  |  |  | regimen was      |
|  |  |  | negatively       |
|  |  |  | associated with  |
|  |  |  | cognitive        |
|  |  |  | impairment in    |
|  |  |  | both colon       |

|        |            |                  |              |              |            | cancer and         |
|--------|------------|------------------|--------------|--------------|------------|--------------------|
|        |            |                  |              |              |            | rectal cancer.     |
|        |            |                  |              |              |            | - Chemotherapy and |
|        |            |                  |              |              |            | radiotherapy       |
|        |            |                  |              |              |            | did not impose     |
|        |            |                  |              |              |            | marked adverse     |
|        |            |                  |              |              |            | cognitive          |
|        |            |                  |              |              |            | effects in         |
|        |            |                  |              |              |            | colorectal         |
|        |            |                  |              |              |            | cancer             |
|        |            |                  |              |              |            | patients.          |
| Lee e  | Longitudin | Stage II and III | Received the | Before and   | Objective  | OCI: peripheral    |
| al.    | al study   | colon cancer     | combination  | after 6      | scale: not | neuropathy         |
| (2016) |            | (n=56)           | adjuvant     | cycles of    | report     | developed in 31    |
| (2010) |            | Mean age 59.5    | chemotherapy | adjuvant     | Subjective | patients (55.4%)   |
| Korea  |            | Education level: | FOLFOX       | chemotherapy | scale:     | after oxaliplatin  |
|        |            | not report       |              |              | EORTC QLQ- | administration.    |
|        |            |                  |              |              | C30        | SCI: not report    |

|  |  |  | - Patients  | with  |
|--|--|--|-------------|-------|
|  |  |  | peripheral  |       |
|  |  |  | neuropathy  |       |
|  |  |  | scored low  | ı in  |
|  |  |  | emotional   | and   |
|  |  |  | cognitive   |       |
|  |  |  | functioning | •     |
|  |  |  | -Age was    | an    |
|  |  |  | important   |       |
|  |  |  | factor      |       |
|  |  |  | influencing | QoL   |
|  |  |  | during      |       |
|  |  |  | chemotherap | y in  |
|  |  |  | this study  | , as  |
|  |  |  | patients ov | er 70 |
|  |  |  | years       |       |
|  |  |  | experienced |       |
|  |  |  | worsening   | of    |

|        |     |                  |              |             |            | their gen    | eral |
|--------|-----|------------------|--------------|-------------|------------|--------------|------|
|        |     |                  |              |             |            | health statu | ıs.  |
|        |     |                  |              |             |            | -There was   | a    |
|        |     |                  |              |             |            | decrease     | in   |
|        |     |                  |              |             |            | emotional    | and  |
|        |     |                  |              |             |            | cognitive    |      |
|        |     |                  |              |             |            | functions.   |      |
| Lin et | RCT | Stage I to III   | After        | At baseline | Objective  | OCI/SCI:     | not  |
| al.    |     | CRC (n=60)       | radical      | 3 mo        | scale: not | report       |      |
| (2022) |     | Experimental     | resection    | 6 mo        | report     | - Among      | the  |
| (2022) |     | group(n=27): CBT | and          |             | Subjective | cognitive    |      |
| China  |     | combined with    | chemotherapy |             | scale:     | status       |      |
|        |     | exercise         |              |             | FACT-Cog   | dimensions,  |      |
|        |     | intervention for |              |             |            | others       |      |
|        |     | 6 mo             |              |             |            | evaluation   |      |
|        |     | Mean age 52      |              |             |            | score was    | the  |
|        |     | Median education |              |             |            | highest,     |      |
|        |     | years 12         |              |             |            | followed     | by   |
|        |     |                  |              |             |            | corrected    |      |

| Control          |  | cognitive   | )        |
|------------------|--|-------------|----------|
| group(n=28): no  |  | impairmer   | nt,      |
| exercise         |  | cognitive   | <b>5</b> |
| intervention     |  | ability,    | and      |
| Mean age 51      |  | impact      | on       |
| Median education |  | quality c   | of life. |
| years 12         |  | -To less ex | ercise,  |
|                  |  | bed         | rest,    |
|                  |  | chemother   | capy,    |
|                  |  | and         | other    |
|                  |  | internal    | and      |
|                  |  | external    |          |
|                  |  | factors,    |          |
|                  |  | patients    | often    |
|                  |  | appear      |          |
|                  |  | anxiety,    |          |
|                  |  | depression  | on, and  |
|                  |  | other n     |          |
|                  |  | emotions    |          |
|                  |  | emotions    | and      |

|  |  |  | varying degrees     |
|--|--|--|---------------------|
|  |  |  | of cognitive        |
|  |  |  | impairment.         |
|  |  |  | - Cognitive         |
|  |  |  | impairment and      |
|  |  |  | the scores of       |
|  |  |  | each dimension      |
|  |  |  | were positively     |
|  |  |  | correlated with     |
|  |  |  | quality of life.    |
|  |  |  | - CBT combined with |
|  |  |  | body-building       |
|  |  |  | Baduanjin           |
|  |  |  | exercise can        |
|  |  |  | improve CRF and     |
|  |  |  | cognitive           |
|  |  |  | impairment in       |
|  |  |  | CRC patients        |
|  |  |  | after               |

|        |           |                |              |               |            | chemotherapy,   |
|--------|-----------|----------------|--------------|---------------|------------|-----------------|
|        |           |                |              |               |            | and improve     |
|        |           |                |              |               |            | their QOL.      |
| Liu et | Cross-    | Stage I to IV  | Undergone    | Only for once | Objective  | OCI/SCI: not    |
| al.    | sectional | CRC (n=29)     | chemotherapy |               | scale:     | report          |
| (2022) | study     | Mean age       |              |               | MMSE, MoCA | - Patients had  |
| (2022) |           | 58.21(8.55)    |              |               | Subjective | decreased       |
| UK     |           | Mean education |              |               | scale:     | scores of MMSE, |
|        |           | years          |              |               | FACT-Cog   | MoCA and FACT-  |
|        |           | 14.03(1.57)    |              |               |            | Cog when        |
|        |           | HCs(n=29)      |              |               |            | compared with   |
|        |           | Mean age       |              |               |            | those of HCs.   |
|        |           | 56.97(7.02)    |              |               |            | - Patients      |
|        |           | Mean education |              |               |            | presented with  |
|        |           | years          |              |               |            | decreased       |
|        |           | 14.62(1.45)    |              |               |            | scores of       |
|        |           |                |              |               |            | cognitive       |
|        |           |                |              |               |            | function        |
|        |           |                |              |               |            | assessment,     |

|  |  |  | which suggest    | ted |
|--|--|--|------------------|-----|
|  |  |  | that             |     |
|  |  |  | chemotherapy     |     |
|  |  |  | might le         | ead |
|  |  |  | cognitive        |     |
|  |  |  | dysfunction.     |     |
|  |  |  | - CRC patients h | nad |
|  |  |  | decreased        |     |
|  |  |  | scores of MMS    | SE, |
|  |  |  | MoCA and FAC     |     |
|  |  |  | Cog, whi         |     |
|  |  |  | implied th       |     |
|  |  |  | these patier     |     |
|  |  |  | following        | 105 |
|  |  |  |                  |     |
|  |  |  | chemotherapy     | ,   |
|  |  |  | had impair       | red |
|  |  |  | cognitive        |     |
|  |  |  | function.        |     |

|          |            |                  |              |              |            | - Cognitive         |
|----------|------------|------------------|--------------|--------------|------------|---------------------|
|          |            |                  |              |              |            | impairment was      |
|          |            |                  |              |              |            | observed in CRC     |
|          |            |                  |              |              |            | patients            |
|          |            |                  |              |              |            | following           |
|          |            |                  |              |              |            | chemotherapy.       |
| Sales et | Longitudin | Stage II and III | Chemotherapy | T1: baseline | Objective  | OCI/SCI: not        |
| al.      | al study   | CRC (n=137)      | and          | assessment   | scale: NP  | report              |
| (2018)   |            | Mean age         | nonchemother | at 1 to 3    | tests      | - Patients with CRC |
|          |            | 62.5(9.4)        | ару          | months after | Subjective | who received        |
| Brazil   |            | Mean education   |              | surgery and  | scale: the | adjuvant 5-         |
|          |            | years 7.6(3.7)   |              | before the   | Everyday   | fluorouracil        |
|          |            |                  |              | start of     | Cognition  | with or without     |
|          |            |                  |              | adjuvant     | questionna | oxaliplatin         |
|          |            |                  |              | therapy.     | ire        | presented with a    |
|          |            |                  |              | T2: follow-  |            | decline in          |
|          |            |                  |              | up           |            | executive           |
|          |            |                  |              | assessment   |            | function after      |
|          |            |                  |              | for the CTh+ |            | 12 months           |

|         |           |                  |              | patients      |             | compared with    |
|---------|-----------|------------------|--------------|---------------|-------------|------------------|
|         |           |                  |              | were          |             | patients with    |
|         |           |                  |              | conducted 12  |             | localized        |
|         |           |                  |              | months after  |             | disease who had  |
|         |           |                  |              | chemotherapy  |             | not received     |
|         |           |                  |              | initiation.   |             | chemotherapy.    |
| Tong et | Cross-    | CRC (n=98)       | After        | Only for once | Objective   | OCI/SCI: not     |
| al.     | sectional | Study group      | chemotherapy |               | scale:      | report           |
|         | study     | (n=98)           |              |               | WAIS-RC,    | - Attention      |
| (2015)  |           | Mean age 57. 55  |              |               | CMS, CPT,   | dysfunction      |
| China   |           | (10.61)          |              |               | WCST, TMT-A | exists in rectal |
|         |           | Mean education   |              |               | and B,      | cancer patients  |
|         |           | years 5.         |              |               | Halstead    | undergoing       |
|         |           | 83(2.78)         |              |               | Reitan Test | postoperative    |
|         |           | Health controls  |              |               | Battery,    | chemotherapy.    |
|         |           | group(n=48) Mean |              |               |             | -The results     |
|         |           | age 56.30(9.63)  |              |               |             | showed that the  |
|         |           | Mean education   |              |               |             | instantaneous    |
|         |           | years 6.53(3.53) |              |               |             | memory           |

|  |  |  | performance   | of  |
|--|--|--|---------------|-----|
|  |  |  | the study gr  | oup |
|  |  |  | decreased in  |     |
|  |  |  | early stage   |     |
|  |  |  |               |     |
|  |  |  | chemotherapy, | ,   |
|  |  |  | but           | the |
|  |  |  | difference    | was |
|  |  |  | not           |     |
|  |  |  | statistically | Y   |
|  |  |  | significant.  |     |
|  |  |  | -The results  | of  |
|  |  |  | meaningless   |     |
|  |  |  | picture       |     |
|  |  |  | recognition   | and |
|  |  |  | associative   |     |
|  |  |  | learning, wh  | ich |
|  |  |  | reflected     |     |
|  |  |  | short-term    |     |
|  |  |  | memory,       | are |

|          |            |                  |              |          |             | lower than those  |
|----------|------------|------------------|--------------|----------|-------------|-------------------|
|          |            |                  |              |          |             | of the control    |
|          |            |                  |              |          |             | group,            |
|          |            |                  |              |          |             | suggesting that   |
|          |            |                  |              |          |             | there is a more   |
|          |            |                  |              |          |             | extensive         |
|          |            |                  |              |          |             | memory            |
|          |            |                  |              |          |             | impairment in     |
|          |            |                  |              |          |             | the study group   |
|          |            |                  |              |          |             | after             |
|          |            |                  |              |          |             | chemotherapy.     |
| Vardy et | Longitudin | Stage I to III   | After        | Baseline | Objective   | OCI: 45% of       |
| al.      | al study   | CRC (n=435)      | surgery and  |          | scale: NP   | patients with     |
| (2014)   |            | Group 1 (n=291): | before any   |          | tests,      | localized CRC had |
|          |            | cognitive        | adjuvant     |          | CANTAB, SET | impairment on NP  |
| Australi |            | function and     | chemotherapy |          | Subjective  | tests and 30% on  |
| a        |            | fatigue in       |              |          | scale:      | CANTAB.           |
|          |            | patients with    |              |          | FACT-Cog    | SCI: perceived    |
|          |            | localized CRC    |              |          | version 2   | cognitive         |

| Mean age 58.6    |  | impairment rates  |
|------------------|--|-------------------|
| years            |  | were 21% in group |
| Mean education   |  | 1, 18.5% in group |
| years 13.8 (3.4) |  | 2.                |
| Group 2 (n=72):  |  | -Only a weak      |
| patients with    |  | association was   |
| limited          |  | observed          |
| metastatic or    |  | between           |
| locally          |  | cognitive         |
| recurrent CRC    |  | symptoms and      |
| before           |  | neuropsychologi   |
| chemotherapy     |  | cal performance   |
| Mean age 56.9    |  | by GDS on NP      |
| Mean education   |  | tests or CANTAB.  |
| years 13.7 (3.4) |  | -A weak           |
| HC (n=72):       |  | association       |
| comparisons      |  | between OCI and   |
| Mean age 56.2    |  | SCI.              |
|                  |  |                   |

| Mean education   |  | - More women | had    |
|------------------|--|--------------|--------|
| years 13.6 (2.9) |  | cognitive    |        |
|                  |  | impairment   | on     |
|                  |  | the clinica  | al GDS |
|                  |  | than men.    |        |
|                  |  | -Women rep   | orted  |
|                  |  | significant  | tly    |
|                  |  | more fa      | atigue |
|                  |  | than men     | after  |
|                  |  | adjusting    | for    |
|                  |  | group.       |        |
|                  |  | - The        | main   |
|                  |  | cognitive    |        |
|                  |  | domains      |        |
|                  |  | affected     | were   |
|                  |  | complex      |        |
|                  |  | information  | n      |
|                  |  | processing   |        |
|                  |  | speed, aud   | litory |

|          |            |                 |              |              |             | working memory,    |
|----------|------------|-----------------|--------------|--------------|-------------|--------------------|
|          |            |                 |              |              |             | and verbal         |
|          |            |                 |              |              |             | learning           |
|          |            |                 |              |              |             | efficiency.        |
|          |            |                 |              |              |             | - Self-reported    |
|          |            |                 |              |              |             | cognitive          |
|          |            |                 |              |              |             | impairment was     |
|          |            |                 |              |              |             | associated with    |
|          |            |                 |              |              |             | fatigue, poorer    |
|          |            |                 |              |              |             | QOL, and anxiety   |
|          |            |                 |              |              |             | and depression.    |
| Vardy et | Longitudin | Stage III or    | Surgery      | At baseline  | Objective   | OCI: 20% of        |
| al.      | al study   | high-risk stage | and/or       | (before      | scale: NP   | patients with      |
| (2015)   |            | II CRC (n=434)  | chemotherapy | chemotherapy | tests,      | localized CRC had  |
|          |            | Treated with    |              | , if given)  | CANTAB, SET | significant        |
| Australi |            | surgery and     |              | At 6 months  | Subjective  | decline in         |
| a        |            | adjuvant or     |              | At 12 months | scale:      | cognitive function |
|          |            | neoadjuvant     |              | At 24 months | FACT-Cog    | from baseline to   |
|          |            |                 |              |              | version 2   | 12 months.         |

| chemotherapy     |  | OCI: rat    | tes                | of  |
|------------------|--|-------------|--------------------|-----|
| (n=173)          |  | cognitive   |                    |     |
| Median age 57.0  |  | impairment  |                    | for |
| Mean education   |  | patients    | W                  | ith |
| years 13.8 (3.3) |  | localized   | (                  | CRC |
| Patients who     |  | ranged fro  | m 36%              | to  |
| underwent        |  | 52%         | betw               | een |
| surgery but did  |  | baseline    | and                | 24  |
| not receive      |  | months.     |                    |     |
| chemotherapy(n=1 |  | SCI: p      | ercei <sup>.</sup> | ved |
| 16)              |  | cognitive   |                    |     |
| Median age 60.5  |  | impairment  |                    | was |
| Mean education   |  | more comm   | on at              | 6   |
| years 13.7 (3.5) |  | months      |                    | in  |
| Metastatic CRC   |  | participan  | ts                 | who |
| (n=73)           |  | received    |                    |     |
| Median age 55.5  |  | chemothera  | .py (3             | 2%) |
| Mean education   |  | than in the | hose               | who |
| years 13.7 (3.4) |  | did not.    |                    |     |

| HCs (n=72)       | OPG had been been |
|------------------|-------------------|
| ncs (11-72)      | - CRC had impaire |
| Median age 58.5  | attention,        |
| Mean education   | working memory    |
| years 13.6 (2.9) | verbal            |
|                  | learning, ar      |
|                  | processing        |
|                  | speed domains.    |
|                  | - There was a wea |
|                  | association       |
|                  | between           |
|                  | objective         |
|                  | neuropsycholog.   |
|                  | cal performance   |
|                  | and self          |
|                  | reported          |
|                  | cognitive         |
|                  | function.         |
|                  | - A moderat       |
|                  | association wa    |

|  |  |  | found at e    | ach |
|--|--|--|---------------|-----|
|  |  |  | assessment    |     |
|  |  |  | between       |     |
|  |  |  | cognitive     |     |
|  |  |  | symptoms      | and |
|  |  |  | fatigue, Qo   | OL, |
|  |  |  | and           |     |
|  |  |  | anxiety/depre | ess |
|  |  |  | ion.          |     |
|  |  |  | - Women with  | CRC |
|  |  |  | had me        | ore |
|  |  |  | cognitive     |     |
|  |  |  | impairment t  | han |
|  |  |  | men at e      | ach |
|  |  |  | assessment,   | but |
|  |  |  | men had       | a   |
|  |  |  | greater risk  | of  |
|  |  |  | cognitive     |     |

|          |            |                  |              |             |            | decline over       |
|----------|------------|------------------|--------------|-------------|------------|--------------------|
|          |            |                  |              |             |            | time.              |
| Vardy et | Longitudin | CRC (n=25)       | Received     | At baseline | Objective  | OCI: 4/24 (55%) of |
| al.      | al study   | Median age 72    | chemotherapy | 6 months    | scale: NP  | CRC survivors were |
| (2022)   |            | Mean education   | Survivors    | 12 months   | test, WRAT | impaired on        |
| (2022)   |            | years 13.6 (2.5) |              | 24 months   | Subjective | functional tasks   |
| Australi |            | HCs(n=25)        |              |             | scale:     | compared to 4/25   |
| a        |            | Median age 68    |              |             | FACT-Cog   | (16%) of controls. |
|          |            | Mean education   |              |             | version 3, | SCI: not report    |
|          |            | years 13.8 (3.1) |              |             | EORTC QLQ- | -There was no      |
|          |            |                  |              |             | C30        | significant        |
|          |            |                  |              |             |            | difference in      |
|          |            |                  |              |             |            | cognitive          |
|          |            |                  |              |             |            | scores or          |
|          |            |                  |              |             |            | proportion with    |
|          |            |                  |              |             |            | cognitive          |
|          |            |                  |              |             |            | impairment         |
|          |            |                  |              |             |            | between            |
|          |            |                  |              |             |            | survivors and      |

|  |  |  | controls and no |
|--|--|--|-----------------|
|  |  |  | evidence of     |
|  |  |  | accelerated     |
|  |  |  | ageing in CRC   |
|  |  |  | survivors.      |
|  |  |  |                 |
|  |  |  | -There was no   |
|  |  |  | difference in   |
|  |  |  | cognitive       |
|  |  |  | capacity and    |
|  |  |  | function        |
|  |  |  | between CRC     |
|  |  |  | survivors and   |
|  |  |  | controls 6-12   |
|  |  |  | years after     |
|  |  |  | diagnosis.      |
|  |  |  | -There was no   |
|  |  |  | significant     |
|  |  |  | difference in   |
|  |  |  | cognitive       |

|          |           |                  |              |           |             | symptoms in CRC    |
|----------|-----------|------------------|--------------|-----------|-------------|--------------------|
|          |           |                  |              |           |             | survivors          |
|          |           |                  |              |           |             | compared to        |
|          |           |                  |              |           |             | controls.          |
| Visovatt | Cross-    | Stage I to IV    | Before any   | Once in 6 | Objective   | OCI/SCI: not       |
| i et al. | sectional | CRC (n=50)       | treatment    | months    | scale: NP   | report             |
| (2016)   | study     | Healthy          | After        |           | test (ANT,  | -Individuals with  |
| (2010)   |           | participants     | surgery only |           | DS, TMT,    | CRC performed      |
| USA      |           | (n=50)           | Chemotherapy |           | RAVLT)      | worse and          |
|          |           | Mean age 56      | and/or       |           | Subjective  | reported more      |
|          |           | Median education | radiation    |           | scale: AFI, | problems on        |
|          |           | years 16         | therapy      |           | EMQ         | tasks requiring    |
|          |           |                  |              |           |             | attention and      |
|          |           |                  |              |           |             | cognitive          |
|          |           |                  |              |           |             | control.           |
|          |           |                  |              |           |             | - Individuals with |
|          |           |                  |              |           |             | CRC reported       |
|          |           |                  |              |           |             | less               |
|          |           |                  |              |           |             | effectiveness      |

|  |  | with activ  | rities   |
|--|--|-------------|--|
|  |  | requiring   |  |
|  |  | attention   | and  |
|  |  | cognitive   |  |
|  |  | control.    |  |
|  |  | -Older age, | less   |
|  |  | education,  | and  |
|  |  | fatigue     | can  |
|  |  | increase    | risk   |
|  |  | for         | worse  |
|  |  | cognitive   |  |
|  |  | performanc  | e and  |
|  |  | self-repor  | ted  |
|  |  | cognition.  |  |
|  |  | -Objective  |  |
|  |  | measures    | are  |
|  |  | less sens   | sitive   |
|  |  | to          | subtle   |
|  |  | deficits    | in   |
|  |  |             | attention cognitive control.  -Older age, education, fatigue increase for cognitive performanc self-repor cognition.  -Objective measures less sens to |

|          |         |                |               |               |            | attentional      |
|----------|---------|----------------|---------------|---------------|------------|------------------|
|          |         |                |               |               |            | capacity and     |
|          |         |                |               |               |            | cognitive        |
|          |         |                |               |               |            | control          |
|          |         |                |               |               |            | perceived by the |
|          |         |                |               |               |            | individual or    |
|          |         |                |               |               |            | that subtle      |
|          |         |                |               |               |            | changes in       |
|          |         |                |               |               |            | fatigue may not  |
|          |         |                |               |               |            | affect           |
|          |         |                |               |               |            | objective test   |
|          |         |                |               |               |            | performance.     |
| Wilson   | Case-   | CRC (n= 88)    | Chemotherapy  | Only for once | Objective  | OCI/SCI: not     |
| et al.   | control | Mean age       | , surgery, or |               | scale: not | report           |
| (2017)   | study   | 65.9(9.7)      | radiation     |               | report     | -Survivors did   |
|          |         | Spouses (n=40) | therapy       |               | Subjective | experience mild  |
| Australi |         | Mean age       |               |               | scale:     | levels of        |
| a        |         | 66.0(9.1)      |               |               | FACT-Cog   | cognitive        |
|          |         |                |               |               | version 3  | decline          |

| Education level: |  |     | accompanied   | by   |
|------------------|--|-----|---------------|------|
| not report       |  |     | depression    |      |
|                  |  |     | following     |      |
|                  |  |     | treatment.    |      |
|                  |  | - 1 | Neither inter | cnal |
|                  |  |     | locus of cont | crol |
|                  |  |     | nor optimism  | in   |
|                  |  |     | colorectal    |      |
|                  |  |     | cancer        |      |
|                  |  |     | survivors     |      |
|                  |  |     | influences    |      |
|                  |  |     | correlation   |      |
|                  |  |     | between       |      |
|                  |  |     | cognition     | and  |
|                  |  |     | depression.   |      |
|                  |  | - : | Health c      | care |
|                  |  |     | providers     |      |
|                  |  |     | should r      | note |
|                  |  |     | individual    |      |

|  |  |  | differences i      | in  |
|--|--|--|--------------------|-----|
|  |  |  | responses t        | .0  |
|  |  |  | treatment and b    | ne. |
|  |  |  |                    |     |
|  |  |  | alert to th        | ne  |
|  |  |  | impact c           | of  |
|  |  |  | depression c       | on  |
|  |  |  | perceived          |     |
|  |  |  | everyday           |     |
|  |  |  | functioning.       |     |
|  |  |  | - Depression wa    | as  |
|  |  |  | the mos            | st  |
|  |  |  | important          |     |
|  |  |  | correlate wit      | :h  |
|  |  |  | cognitive          |     |
|  |  |  | functioning.       |     |
|  |  |  | - Internal locus o | of  |
|  |  |  | control coul       | Ld  |
|  |  |  | explain            |     |
|  |  |  | additional         |     |

|  |  |  | variance in     |
|--|--|--|-----------------|
|  |  |  | cognitive       |
|  |  |  | function beyond |
|  |  |  | that accounted  |
|  |  |  | for by          |
|  |  |  | depression.     |
|  |  |  | - Confirmed     |
|  |  |  | moderately      |
|  |  |  | strong          |
|  |  |  | relationships   |
|  |  |  | between         |
|  |  |  | depression      |
|  |  |  | around time of  |
|  |  |  | treatment for   |
|  |  |  | colorectal      |
|  |  |  | cancer and      |
|  |  |  | adverse         |
|  |  |  | cognitive       |
|  |  |  | functioning     |

|         |           |                    |               |               |             | experienced      |
|---------|-----------|--------------------|---------------|---------------|-------------|------------------|
|         |           |                    |               |               |             | following        |
|         |           |                    |               |               |             | treatment.       |
| Yang et | Cross-    | Stage I to III     | New           | Only for once | Objective   | OCI/SCI: not     |
| al.     | sectional | CRC (n=63)         | diagnosis     |               | scale:      | report           |
| (2023)  | study     | Group A $(n = 13)$ | (pretreatmen  |               | MoCA, NP    | -Findings reveal |
| (2023)  |           | Mean age           | t)            |               | tests       | that the three   |
| China   |           | 62.15(9.81)        | 2 years since |               | - Attention | groups           |
|         |           | Mean education     | chemotherapy  |               | :           | exhibited        |
|         |           | years              | completion    |               | - Memory:   | comparable       |
|         |           | 13.46(3.97)        | > 2 years     |               | WMS-III     | subjective       |
|         |           | Group B (n=24)     | since         |               | -Executive  | cognitive        |
|         |           | Mean age           | chemotherapy  |               | function    | functions and    |
|         |           | 57.04(8.42)        | completion    |               | and:        | objective        |
|         |           | Mean education     |               |               | WAIS-IV,    | performances in  |
|         |           | years              |               |               | SCWT        | overall          |
|         |           | 14.17(4.20)        |               |               | Subjective  | cognitive        |
|         |           |                    |               |               | : FACT-Cog  | function and     |

| Group C (n = 26)   |  | episodic   | <u> </u> |
|--------------------|--|------------|----------|
| Mean age           |  | memory.    |          |
| 58.08(7.74)        |  | - Patients | with CRC |
| Mean education     |  | who        | receive  |
| years 13.38        |  | chemothe   |          |
| (3.14);            |  |            |          |
| Matched set (n =   |  |            | high     |
|                    |  | levels     | of       |
| 36)                |  | cognitiv   | re       |
| Group A $(n = 12)$ |  | impairme   | ent      |
| Mean age           |  | within     | 2 years  |
| 61.05(9.94)        |  | of t       | reatment |
| Mean education     |  | completi   | on.      |
| years              |  | - Patients | who      |
| 13.25(4.07)        |  | complete   | d        |
| Group B (n = 12)   |  | chemothe   | rapy in  |
| Mean age           |  | 1 to 2     | 2 years  |
| 54.75(7.93)        |  | (Group     | B) had   |
|                    |  | worse so   | cores in |
|                    |  | cognitiv   | re       |

| Mean education   |  | function           |
|------------------|--|--------------------|
| years            |  | compared with      |
| 14.50(4.08)      |  | the others.        |
| Group C (n = 12) |  | -Noted a dearth of |
| Mean age         |  | studies on the     |
| 59.42(7.42)      |  | evaluation of      |
| Mean education   |  | changes in         |
| years            |  | specific           |
| 14.00(2.66)      |  | cognitive          |
|                  |  | functions at       |
|                  |  | various time       |
|                  |  | points within 1    |
|                  |  | to 2 years of      |
|                  |  | the completion     |
|                  |  | of                 |
|                  |  | chemotherapy.      |
|                  |  | - Adjuvant         |
|                  |  | chemotherapy       |
|                  |  | had                |
|                  |  |                    |

|          |         |                |              |               |             | significar | ntly   |
|----------|---------|----------------|--------------|---------------|-------------|------------|--------|
|          |         |                |              |               |             | deleteriou | ıs     |
|          |         |                |              |               |             | effects    | on     |
|          |         |                |              |               |             | attention  | and    |
|          |         |                |              |               |             | processing | ı      |
|          |         |                |              |               |             | speed      | in     |
|          |         |                |              |               |             | patients   | with   |
|          |         |                |              |               |             | CRC,       | which  |
|          |         |                |              |               |             | impacted   |        |
|          |         |                |              |               |             | cognitive  |        |
|          |         |                |              |               |             | symptoms   |        |
|          |         |                |              |               |             | lasting    | for    |
|          |         |                |              |               |             | approximat | cely   |
|          |         |                |              |               |             | two years. |        |
| Zhang et | Case-   | Stage I to III | Receiving    | Preoperative  | Objective   | OCI: ni    | neteen |
| al.      | control | CRC (total     | surgery and  | ly and at the | scale:      | patients   | were   |
| (2018)   | study   | sample n=77)   | non-         | seventh day   | MMSE, VVLT, | diagnosed  | as     |
|          |         |                | chemotherapy | after         | DST         | cognitive  |        |
| China    |         |                |              | operation     |             |            |        |

| Group cognitive  |   |   | impairment         |
|------------------|---|---|--------------------|
| impairment (n    |   |   | (24.7%).           |
| =58)             |   |   | SCI: not report    |
| Mean age 73.     |   |   | - Diabetes history |
| 30(4.80)         |   |   | fasting over       |
| Mean education   |   |   | three days after   |
| years 5. 80(2.   |   |   | surgery and SIRS   |
| 30)              |   |   | score>3 on the     |
| Group non-       |   |   | second day after   |
| cognitive        |   |   | surgery were       |
| impairment (n =  |   |   | risk factors of    |
| 19) Mean age     |   |   | early cognitive    |
| 69.30(4.60) Mean |   |   | impairment in      |
| education years  |   |   | the elderly        |
| 7.80(2.60)       |   |   | patients after     |
|                  |   |   | radical            |
|                  |   |   | resection for      |
|                  |   |   | CRC.               |
|                  | I | 1 |                    |

| Zhang et | Cohort | Stage II and III | Receive    | One     | day  | Objective   | OCI: A total of 19 |
|----------|--------|------------------|------------|---------|------|-------------|--------------------|
| al.      | study  | CRC              | colorectal | before  |      | scale:      | patients (24.7%)   |
| (2010)   |        | Non-cognitive    | surgery    | Seven   | days | MMSE, VVLT, | were identified as |
| (2019)   |        | impairment (n =  |            | after   |      | DST         | cognitive          |
| China    |        | 58)              |            | surgery |      | Subjective  | impairment.        |
|          |        | Mean age         |            |         |      | scale: not  | SCI: not report    |
|          |        | 69.3(4.6)        |            |         |      | report      | -The test battery  |
|          |        | Mean education   |            |         |      |             | scores in the      |
|          |        | years 7.8 (2.6)  |            |         |      |             | cognitive          |
|          |        | Cognitive        |            |         |      |             | impairment         |
|          |        | impairment (n =  |            |         |      |             | group were         |
|          |        | 19)              |            |         |      |             | significantly      |
|          |        | Mean age         |            |         |      |             | lower than those   |
|          |        | 73.3(4.8)        |            |         |      |             | in the non-        |
|          |        | Mean education   |            |         |      |             | cognitive          |
|          |        | years 5.8 (2.3)  |            |         |      |             | impairment         |
|          |        |                  |            |         |      |             | group.             |
|          |        |                  |            |         |      |             | -There was no      |
|          |        |                  |            |         |      |             | significant        |

|        |          |                  |              |               |             | difference        |
|--------|----------|------------------|--------------|---------------|-------------|-------------------|
|        |          |                  |              |               |             | between the       |
|        |          |                  |              |               |             | cognitive         |
|        |          |                  |              |               |             | impairment and    |
|        |          |                  |              |               |             | non-cognitive     |
|        |          |                  |              |               |             | impairment        |
|        |          |                  |              |               |             | groups in terms   |
|        |          |                  |              |               |             | of gender.        |
| Zhou   | ct Case- | Total sample     | After        | Only for once | Objective   | OCI: 117 patients |
| al.    | control  | CRC (n=386)      | chemotherapy |               | scale: MMSE | with CRC were     |
| (2021) | study    | Mean age         | Survivorship |               | Subjective  | cognitive         |
|        |          | 64.73(10.8)      |              |               | scale: not  | impairment.       |
| China  |          | Median education |              |               | report      | SCI: not report   |
|        |          | level 12         |              |               |             | -This model could |
|        |          |                  |              |               |             | predict           |
|        |          |                  |              |               |             | cognitive         |
|        |          |                  |              |               |             | impairment risk   |
|        |          |                  |              |               |             | in CRC patients   |
|        |          |                  |              |               |             | after             |

|  |  |  | chemotherapy at  |
|--|--|--|------------------|
|  |  |  | an early stage.  |
|  |  |  | - The model      |
|  |  |  | suggested that   |
|  |  |  | advanced age,    |
|  |  |  | colostomy,       |
|  |  |  | diabetes, lower  |
|  |  |  | education, and   |
|  |  |  | later            |
|  |  |  | pathological     |
|  |  |  | stage could be   |
|  |  |  | risk factors for |
|  |  |  | cognitive        |
|  |  |  | impairment in    |
|  |  |  | CRC patients     |
|  |  |  | after            |
|  |  |  | chemotherapy.    |

AFI: Attentional Function Index; ANT: Attention Network Test; CANTAB: Clinical Battery and Computer-Based Cambridge Automated Battery; CBT: Cognitive Behavior Therapy; CDT:

Clock Drawing Test; CMS: Clinical Memory Scale; COWA: the Controlled Oral Word Association of the Multilingual Aphasia Examination; CPT: Continuous Performance Test; CRC: Colorectal Cancer; CRF: Cancer-Related Fatigue; CTh+: receive chemotherapy; CTh-: without an adjuvant chemotherapy; Delayed-Mem: Delayed Memory; Delayed-Mem-Q: Delayed Memory-Questions; DST: Digital Span Test; DSymbol: WAIS-R Digit Symbol; EMQ: Everyday Memory Questionnaire; EORTC QLQ-C30: European Organization for Research and Treatment of Cancer Core Questionnaire; FACT-Coq: Functional Assessment of Cancer Therapy-Coqnitive Function questionnaire; FOLFOX4: Combination of Oxaliplatin and 5-Fluorouracil Plus Leucovorin; GDS: Global Deficit Scores; HCs: Health Controls; HVLT-R: Hopkins Verbal Learning Test-Revised; Imm-Mem: immediate memory; Imm-Mem-Q: immediate memory-questions; LMWT: Luria Memory Words Test; MMSE: Mini Mental State Examination; MoCA: the Montreal Cognitive Assessment; NP tests: the clinic neuropsychological tests; PCI: perceived cognitive impairments; OCI: objective cognitive impairment; QoL: Quality of Life; RAVLT: Rey Auditory Verbal Learning Test; SCI: subjective cognitive impairment; SCWT: Stroop Color and Word Test; SET: Modified Six Elements Test; TMT: Trial Making Test; TMT A and B: Trial Making Test A and B; VVLT: Visual Verbal Learning Test; WAIS-R: Wechsler Adult Intelligence Scale-Revised; WAIS-RC: Wechsler Adult Intelligence Scale Revised in China;

WCST: Wisconsin Card Sorting Test; WRAT: Wide Range Achievement Test; WMS-III: Word Lists
Test from the Wechsler Memory Scale-II