

## SUPPLEMENTAL MATERIAL

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Dear Sir/Madam,

thank you very much for your willingness, time and attention you devote to completing this form. Answering the questions below will provide you with an opportunity to aid a study aiming to determine, or at the very least partially clarify, the cause of chronic lymphocytic leukemia, and at the same time the risks of the disease development prior to its onset.

This study intends to identify the factors leading to the development of chronic lymphocytic leukemia, in particular past illnesses, occupations, lifestyles and medicines used. These factors and their influence on the disease development are still not well known.

Your privacy is important to us and so we would like to assure you that the information you provide in this form will be processed completely anonymously and the confidentiality of any personal data is guaranteed.

Patient identification code: \_\_\_\_\_

The patient was informed about the purpose and nature of this study and agreed with processing his/her data:

☐ Yes ☐ No

Physician providing information:

The patient completed the questionnaire:

☐ on his/her own ☐ in cooperation with a physician ☐ with another person

Notes: \_\_\_\_\_

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Date and signature of the patient:

Date, stamp and signature of the physician

## Answer form

### Social history:

Date of birth: \_\_\_\_\_ Gender: ☐ Man ☐ Woman

Nationality: \_\_\_\_\_

Current occupation: \_\_\_\_\_

Primary lifetime occupation: \_\_\_\_\_

Pensioner: ☐ No ☐ Yes

Marital status: \_\_\_\_\_ Number of children born: \_\_\_\_\_

Do you live alone? ☐ No ☐ Yes

### Personal history: Indicate approximate age or date for each positive choice.

#### Substance abuse:

Smoking: ☐ No ☐ Quit smoking, \_\_\_\_\_ years ago ☐ Yes, number of cigarettes: \_\_\_\_\_ / day

Alcohol: ☐ No ☐ Occasionally ☐ Regularly

### Medical history:

#### Childhood diseases:

|                 |                             |                                    |             |                             |                                    |
|-----------------|-----------------------------|------------------------------------|-------------|-----------------------------|------------------------------------|
| Measles:        | <input type="checkbox"/> No | <input type="checkbox"/> Yes _____ | Mumps:      | <input type="checkbox"/> No | <input type="checkbox"/> Yes _____ |
| Rubella:        | <input type="checkbox"/> No | <input type="checkbox"/> Yes _____ | Chickenpox: | <input type="checkbox"/> No | <input type="checkbox"/> Yes _____ |
| Whooping cough: | <input type="checkbox"/> No | <input type="checkbox"/> Yes _____ | Diphtheria: | <input type="checkbox"/> No | <input type="checkbox"/> Yes _____ |
| Scarlet fever:  | <input type="checkbox"/> No | <input type="checkbox"/> Yes _____ | Pneumonia:  | <input type="checkbox"/> No | <input type="checkbox"/> Yes _____ |

Other: \_\_\_\_\_

#### Heart/ Blood circulation:

High blood pressure: ☐ No ☐ Yes Low blood pressure: ☐ No ☐ Yes

Other: \_\_\_\_\_

#### Respiratory diseases:

|                    |                             |                              |         |                             |                              |
|--------------------|-----------------------------|------------------------------|---------|-----------------------------|------------------------------|
| Pneumonia:         | <input type="checkbox"/> No | <input type="checkbox"/> Yes | COPD:   | <input type="checkbox"/> No | <input type="checkbox"/> Yes |
| Chronic sinusitis: | <input type="checkbox"/> No | <input type="checkbox"/> Yes | Asthma: | <input type="checkbox"/> No | <input type="checkbox"/> Yes |
| Tuberculosis:      | <input type="checkbox"/> No | <input type="checkbox"/> Yes |         |                             |                              |

Other: \_\_\_\_\_

#### Metabolism:

Diabetes mellitus: ☐ No ☐ Yes Type 1/ Type 2  
 High cholesterol: ☐ No ☐ Yes  
 Obesity: ☐ No ☐ Yes (BMI: \_\_\_\_\_ )

Other: \_\_\_\_\_

**Malignant diseases (cancer):**

☐ No ☐ Yes \_\_\_\_\_

Localization: \_\_\_\_\_ Status on admission: \_\_\_\_\_

**Therapy:** Chemotherapy (in infusions or pills): ☐ No ☐ Yes \_\_\_\_\_

Radiation therapy: ☐ No ☐ Yes \_\_\_\_\_

Surgery: ☐ No ☐ Yes \_\_\_\_\_

Targeted therapy (monoclonal antibodies, hormonal therapy) ☐ No ☐ Yes \_\_\_\_\_

**Non-malignant diseases (in particular, tumors with no chemotherapy or radiation therapy):** ☐ No ☐ Yes \_\_\_\_\_

**Autoimmune diseases:**

Systemic lupus erythematosus (Lupus): ☐ No ☐ Yes \_\_\_\_\_

Hashimoto's disease: ☐ No ☐ Yes \_\_\_\_\_

Dermatitis herpetiformis: ☐ No ☐ Yes \_\_\_\_\_

Rheumatoid arthritis: ☐ No ☐ Yes \_\_\_\_\_

Multiple sclerosis: ☐ No ☐ Yes \_\_\_\_\_

Sjogren's syndrome: ☐ No ☐ Yes \_\_\_\_\_

Spondyloarthropathy: ☐ No ☐ Yes \_\_\_\_\_

Dermatomyositis: ☐ No ☐ Yes \_\_\_\_\_

Polymyositis: ☐ No ☐ Yes \_\_\_\_\_

Celiac disease: ☐ No ☐ Yes \_\_\_\_\_

Psoriasis: ☐ No ☐ Yes \_\_\_\_\_

Other: \_\_\_\_\_

**Rheumatic diseases (joints pain, joints swelling, producing antibodies that attack the body's own tissues):** \_\_\_\_\_

**Chronic Infectious Diseases:**

Viral: Hepatitis A: ☐ No ☐ Yes \_\_\_\_\_

Hepatitis B: ☐ No ☐ Yes \_\_\_\_\_

Hepatitis C: ☐ No ☐ Yes \_\_\_\_\_

Shingles (Herpes zoster): ☐ No ☐ Yes \_\_\_\_\_

Cold sores (Herpes simplex): ☐ No ☐ Yes \_\_\_\_\_

Infectious mononucleosis: ☐ No ☐ Yes \_\_\_\_\_

Virus EBV: ☐ No ☐ Yes \_\_\_\_\_

Virus HIV: ☐ No ☐ Yes \_\_\_\_\_

Bacterial: Gastritis, peptic ulcers (*Helicobacter pylori*): ☐ No ☐ Yes \_\_\_\_\_

Campylobacteriosis (*Campylobacter jejuni*): ☐ No ☐ Yes \_\_\_\_\_

Chlamydia infection (*Chlamydia psittaci*): ☐ No ☐ Yes \_\_\_\_\_

Lyme disease (*Borrelia burgdorferi*): ☐ No ☐ Yes \_\_\_\_\_

Other significant infections: \_\_\_\_\_

**Chronic and other significant diseases:** \_\_\_\_\_

**Allergies:** \_\_\_\_\_

#### **Vaccination:**

Measles: ☐ No ☐ Yes Mumps: ☐ No ☐ Yes

Rubella: ☐ No ☐ Yes Whooping cough: ☐ No ☐ Yes

Diphtheria: ☐ No ☐ Yes Scarlet fever: ☐ No ☐ Yes

Polio: ☐ No ☐ Yes Tetanus: ☐ No ☐ Yes

Other: \_\_\_\_\_

#### **Surgeries:**

Tonsillectomy: ☐ No ☐ Yes \_\_\_\_\_

Adenoidectomy: ☐ No ☐ Yes \_\_\_\_\_

Appendectomy: ☐ No ☐ Yes \_\_\_\_\_

Cholecystectomy: ☐ No ☐ Yes \_\_\_\_\_

Other: \_\_\_\_\_

#### **Family diseases:**

##### **The occurrence of cancer in your family:**

Mother: ☐ No ☐ Yes \_\_\_\_\_

Father: ☐ No ☐ Yes \_\_\_\_\_

Sibling 1: ☐ No ☐ Yes \_\_\_\_\_

Sibling 2: ☐ No ☐ Yes \_\_\_\_\_

Sibling 3: ☐ No ☐ Yes \_\_\_\_\_

Child 1: ☐ No ☐ Yes \_\_\_\_\_

Child 2:    ☐ No    ☐ Yes \_\_\_\_\_

Child 3:    ☐ No    ☐ Yes \_\_\_\_\_

**The occurrence of autoimmune diseases in your family:**

Mother:    ☐ No    ☐ Yes \_\_\_\_\_

Father:    ☐ No    ☐ Yes \_\_\_\_\_

Sibling 1:    ☐ No    ☐ Yes \_\_\_\_\_

Sibling 2:    ☐ No    ☐ Yes \_\_\_\_\_

Sibling 3:    ☐ No    ☐ Yes \_\_\_\_\_

Child 1:    ☐ No    ☐ Yes \_\_\_\_\_

Child 2:    ☐ No    ☐ Yes \_\_\_\_\_

Child 3:    ☐ No    ☐ Yes \_\_\_\_\_

**The occurrence of chronic or other significant diseases:**

Mother:    ☐ No    ☐ Yes \_\_\_\_\_

Father:    ☐ No    ☐ Yes \_\_\_\_\_

Sibling 1:    ☐ No    ☐ Yes \_\_\_\_\_

Sibling 2:    ☐ No    ☐ Yes \_\_\_\_\_

Sibling 3:    ☐ No    ☐ Yes \_\_\_\_\_

Child 1:    ☐ No    ☐ Yes \_\_\_\_\_

Child 2:    ☐ No    ☐ Yes \_\_\_\_\_

Child 3:    ☐ No    ☐ Yes \_\_\_\_\_

|    | A                             | B   | X                | Y                                     | Z                  | AA                             | AB                | AC                               | AD                      | AE                                     | AF                   | AG              | AH              | AI              | AJ                         | AK                          | AL                      | AM                  | AN                  | AO                                        | AP                           |
|----|-------------------------------|-----|------------------|---------------------------------------|--------------------|--------------------------------|-------------------|----------------------------------|-------------------------|----------------------------------------|----------------------|-----------------|-----------------|-----------------|----------------------------|-----------------------------|-------------------------|---------------------|---------------------|-------------------------------------------|------------------------------|
| 1  | <b>Supplementary Table 1:</b> |     |                  |                                       |                    |                                |                   |                                  |                         |                                        |                      |                 |                 |                 |                            |                             |                         |                     |                     |                                           |                              |
| 2  | Output from questionnaire     |     |                  |                                       |                    |                                |                   |                                  |                         |                                        |                      |                 |                 |                 |                            |                             |                         |                     |                     |                                           |                              |
| 3  |                               |     |                  |                                       |                    |                                |                   |                                  |                         |                                        |                      |                 |                 |                 |                            |                             |                         |                     |                     |                                           |                              |
| 4  | groups tested                 | N   | Asthma<br>no/yes | Respiratory<br>system - all<br>no/yes | Diabetes<br>no/yes | Hypercholesterolemia<br>no/yes | Obesity<br>no/yes | Other malignant<br>tumors no/yes | Benign<br>tumors no/yes | Autoimmune<br>diseases - all<br>no/yes | Rheumatism<br>no/yes | Hep A<br>no/yes | Hep B<br>no/yes | Hep C<br>no/yes | Herpes<br>zoster<br>no/yes | Herpes<br>simplex<br>no/yes | Mononucleosis<br>no/yes | EBV virus<br>no/yes | HIV virus<br>no/yes | Gastritis,<br>stomach<br>ulcers<br>no/yes | Campylobacteriosis<br>no/yes |
| 5  | all                           | 573 | 542/31           | 355/218                               | 465/108            | 365/208                        | 433/140           | 488/85                           | 461/112                 | 506/67                                 | 437/136              | 518/55          | 561/12          | 571/2           | 466/107                    | 391/182                     | 555/18                  | 565/8               | 573/0               | 517/56                                    | 570/3                        |
| 6  | mono                          | 518 | 489/29           | 319/199                               | 421/97             | 340/178                        | 393/125           | 442/76                           | 417/101                 | 456/62                                 | 393/125              | 465/53          | 506/12          | 516/2           | 426/92                     | 361/157                     | 501/17                  | 511/7               | 518/0               | 464/54                                    | 516/2                        |
| 7  | oligo                         | 55  | 53/2             | 36/19                                 | 44/11              | 25/30                          | 40/15             | 46/9                             | 44/11                   | 50/5                                   | 44/11                | 53/2            | 55/0            | 55/0            | 40/15                      | 30/25                       | 54/1                    | 54/1                | 55/0                | 53/2                                      | 54/1                         |
| 8  | monoU                         | 279 | 264/15           | 173/106                               | 226/53             | 191/88                         | 212/67            | 234/45                           | 223/56                  | 239/40                                 | 209/70               | 252/27          | 272/7           | 278/1           | 230/49                     | 200/79                      | 270/9                   | 274/5               | 279/0               | 249/30                                    | 277/2                        |
| 9  | monoM                         | 239 | 225/14           | 146/93                                | 195/44             | 149/90                         | 181/58            | 208/31                           | 194/45                  | 217/22                                 | 184/55               | 213/26          | 234/5           | 238/1           | 196/43                     | 161/78                      | 231/8                   | 237/2               | 239/0               | 215/24                                    | 239/0                        |
| 10 | oligoU                        | 16  | 15/1             | 10/6                                  | 12/4               | 10/6                           | 13/3              | 12/4                             | 14/2                    | 15/1                                   | 15/1                 | 16/0            | 16/0            | 16/0            | 11/5                       | 10/6                        | 16/0                    | 16/0                | 16/0                | 15/1                                      | 16/0                         |
| 11 | oligoD                        | 21  | 21/0             | 13/8                                  | 18/3               | 5/16                           | 17/4              | 17/4                             | 15/6                    | 19/2                                   | 15/6                 | 21/0            | 21/0            | 21/0            | 14/7                       | 10/11                       | 21/0                    | 21/0                | 21/0                | 21/0                                      | 21/0                         |
| 12 | oligoM                        | 18  | 17/1             | 13/5                                  | 14/4               | 10/8                           | 10/8              | 17/1                             | 15/3                    | 16/2                                   | 14/4                 | 16/2            | 18/0            | 18/0            | 15/3                       | 10/8                        | 17/1                    | 17/1                | 18/0                | 17/1                                      | 17/1                         |
| 13 | s1                            | 19  | 18/1             | 9/10                                  | 18/1               | 14/5                           | 15/4              | 15/4                             | 15/4                    | 16/3                                   | 15/4                 | 15/4            | 19/0            | 19/0            | 16/3                       | 12/7                        | 18/1                    | 18/1                | 19/0                | 13/6                                      | 19/0                         |
| 14 | s2                            | 18  | 17/1             | 13/5                                  | 13/5               | 12/6                           | 16/2              | 15/3                             | 15/3                    | 17/1                                   | 13/5                 | 16/2            | 18/0            | 18/0            | 14/4                       | 13/5                        | 17/1                    | 17/1                | 18/0                | 16/2                                      | 18/0                         |
| 15 | s3                            | 4   | 3/1              | 2/2                                   | 4/0                | 4/0                            | 4/0               | 2/2                              | 4/0                     | 2/2                                    | 3/1                  | 4/0             | 4/0             | 4/0             | 3/1                        | 4/0                         | 4/0                     | 4/0                 | 4/0                 | 4/0                                       | 4/0                          |
| 16 | s14                           | 4   | 3/1              | 1/3                                   | 3/1                | 2/2                            | 3/1               | 4/0                              | 2/2                     | 4/0                                    | 3/1                  | 4/0             | 4/0             | 4/0             | 4/0                        | 3/1                         | 4/0                     | 4/0                 | 4/0                 | 4/0                                       | 4/0                          |
| 17 | s99                           | 4   | 4/0              | 4/0                                   | 4/0                | 2/2                            | 3/1               | 4/0                              | 3/1                     | 3/1                                    | 1/3                  | 4/0             | 4/0             | 4/0             | 2/2                        | 2/2                         | 4/0                     | 4/0                 | 4/0                 | 4/0                                       | 4/0                          |
| 18 | s202                          | 5   | 4/1              | 2/3                                   | 4/1                | 1/4                            | 4/1               | 4/1                              | 5/0                     | 3/2                                    | 3/2                  | 5/0             | 5/0             | 5/0             | 3/2                        | 3/2                         | 5/0                     | 5/0                 | 5/0                 | 5/0                                       | 5/0                          |

|    | A                         | B   | AQ                             | AR              | AS               | AT                       | AU                                  | AV                        | AW                            | AX                                            | AY                                               | AZ                        | BA                                 | BB                                   | BC                                                   | BD                                | BE                                      | BF                                               |
|----|---------------------------|-----|--------------------------------|-----------------|------------------|--------------------------|-------------------------------------|---------------------------|-------------------------------|-----------------------------------------------|--------------------------------------------------|---------------------------|------------------------------------|--------------------------------------|------------------------------------------------------|-----------------------------------|-----------------------------------------|--------------------------------------------------|
| 1  | Supplementary Table 1:    |     |                                |                 |                  |                          |                                     |                           |                               |                                               |                                                  |                           |                                    |                                      |                                                      |                                   |                                         |                                                  |
| 2  | Output from questionnaire |     |                                |                 |                  |                          |                                     |                           |                               |                                               |                                                  |                           |                                    |                                      |                                                      |                                   |                                         |                                                  |
| 3  |                           |     |                                |                 |                  |                          |                                     |                           |                               |                                               |                                                  |                           |                                    |                                      |                                                      |                                   |                                         |                                                  |
| 4  | groups tested             | N   | Chlamydia infections<br>no/yes | Borrelia no/yes | Allergies no/yes | Tonsils operation no/yes | Pharyngeal tonsils operation no/yes | Appendix operation no/yes | Gall bladder operation no/yes | Tonsils / pharyngeal tonsils operation no/yes | (Pharyngeal) Tonsils / appendix operation no/yes | Parents with tumor no/yes | Parents/siblings with tumor no/yes | Tumor in 1st degree relatives no/yes | Hematological disease in 1st degree relatives no/yes | CI in 1st degree relatives no/yes | Parents with autoimmune disorder no/yes | Parents/siblings with autoimmune disorder no/yes |
| 5  | all                       | 573 | 559/14                         | 509/64          | 427/146          | 471/102                  | 484/89                              | 462/111                   | 483/90                        | 423/150                                       | 339/234                                          | 327/246                   | 271/302                            | 256/317                              | 532/41                                               | 561/12                            | 449/124                                 | 346/137                                          |
| 6  | mono                      | 518 | 504/14                         | 461/57          | 388/130          | 425/93                   | 436/82                              | 417/101                   | 438/80                        | 380/138                                       | 304/214                                          | 300/218                   | 252/266                            | 239/279                              | 484/34                                               | 510/8                             | 407/111                                 | 395/123                                          |
| 7  | oligo                     | 55  | 55/0                           | 48/7            | 39/16            | 46/9                     | 48/7                                | 45/10                     | 45/10                         | 43/12                                         | 35/20                                            | 27/28                     | 19/36                              | 17/38                                | 48/7                                                 | 51/4                              | 42/13                                   | 41/14                                            |
| 8  | monoU                     | 279 | 272/7                          | 250/29          | 212/67           | 230/49                   | 235/44                              | 229/50                    | 239/40                        | 206/73                                        | 166/113                                          | 164/115                   | 134/145                            | 125/154                              | 261/18                                               | 276/3                             | 212/67                                  | 206/73                                           |
| 9  | monoM                     | 239 | 232/7                          | 211/28          | 176/63           | 195/44                   | 201/38                              | 188/51                    | 199/40                        | 174/65                                        | 138/101                                          | 136/103                   | 118/121                            | 114/125                              | 223/16                                               | 234/5                             | 195/44                                  | 189/50                                           |
| 10 | oligoU                    | 16  | 16/0                           | 16/0            | 10/6             | 14/2                     | 14/2                                | 15/1                      | 14/2                          | 13/3                                          | 12/4                                             | 8/8                       | 5/11                               | 4/12                                 | 14/2                                                 | 15/1                              | 13/3                                    | 13/3                                             |
| 11 | oligoD                    | 21  | 21/0                           | 18/3            | 17/4             | 17/4                     | 19/2                                | 17/4                      | 19/2                          | 17/4                                          | 13/8                                             | 11/10                     | 8/13                               | 8/13                                 | 20/1                                                 | 20/1                              | 15/6                                    | 15/6                                             |
| 12 | oligoM                    | 18  | 18/0                           | 14/4            | 12/6             | 15/3                     | 15/3                                | 13/5                      | 12/6                          | 13/5                                          | 10/8                                             | 8/10                      | 6/12                               | 5/13                                 | 14/4                                                 | 16/2                              | 14/4                                    | 13/5                                             |
| 13 | s1                        | 19  | 19/0                           | 17/2            | 15/4             | 15/4                     | 17/2                                | 18/1                      | 19/0                          | 14/5                                          | 14/5                                             | 10/9                      | 9/10                               | 7/12                                 | 19/0                                                 | 19/0                              | 13/6                                    | 13/6                                             |
| 14 | s2                        | 18  | 18/0                           | 15/3            | 12/6             | 15/3                     | 15/3                                | 15/3                      | 12/6                          | 13/5                                          | 10/8                                             | 12/6                      | 11/7                               | 10/8                                 | 16/2                                                 | 16/2                              | 16/2                                    | 15/3                                             |
| 15 | s3                        | 4   | 4/0                            | 4/0             | 3/1              | 3/1                      | 3/1                                 | 3/1                       | 4/0                           | 3/1                                           | 2/2                                              | 2/2                       | 2/2                                | 1/3                                  | 4/0                                                  | 4/0                               | 4/0                                     | 4/0                                              |
| 16 | s14                       | 4   | 4/0                            | 3/1             | 3/1              | 2/2                      | 3/1                                 | 2/2                       | 3/1                           | 2/2                                           | 0/4                                              | 4/0                       | 3/1                                | 3/1                                  | 4/0                                                  | 4/0                               | 3/1                                     | 3/1                                              |
| 17 | s99                       | 4   | 4/0                            | 3/1             | 3/1              | 2/2                      | 3/1                                 | 3/1                       | 2/2                           | 2/2                                           | 2/2                                              | 3/1                       | 1/3                                | 0/4                                  | 4/0                                                  | 4/0                               | 4/0                                     | 4/0                                              |
| 18 | s202                      | 5   | 5/0                            | 5/0             | 4/1              | 4/1                      | 5/0                                 | 3/2                       | 4/1                           | 4/1                                           | 2/3                                              | 4/1                       | 3/2                                | 3/2                                  | 5/0                                                  | 5/0                               | 5/0                                     | 5/0                                              |

**Supplementary Table 1:**  
Output from questionnaire

| groups tested | N   | Autoimmune disorder in 1st degree relatives no/yes | Questionnaire before treatment no/yes/NA | Median follow-up (range) [mo] |
|---------------|-----|----------------------------------------------------|------------------------------------------|-------------------------------|
| all           | 573 | 416/157                                            | 229/322/22                               | 62.8<br>(1.6-298.9)           |
| mono          | 518 | 376/142                                            | 312/286/19                               | 62.7<br>(1.6-298.9)           |
| oligo         | 55  | 40/15                                              | 16/36/3                                  | 62.9<br>(8.4-210.0)           |
| monoU         | 279 | 193/86                                             | 152/118/9                                | 62.1<br>(3.5-274.1)           |
| monoM         | 239 | 183/56                                             | 61/168/10                                | 64.7<br>(1.6-298.9)           |
| oligoU        | 16  | 13/3                                               | 6/9/1                                    | 45.0<br>(8.4-154.1)           |
| oligoD        | 21  | 15/6                                               | 6/14/1                                   | 50.1<br>(8.9-210.0)           |
| oligoM        | 18  | 12/6                                               | 4/13/1                                   | 78.3<br>(20.0-141.2)          |
| s1            | 19  | 12/7                                               | 10/7/2                                   | 54.1<br>(12.2-184.2)          |
| s2            | 18  | 15/3                                               | 8/8/2                                    | 44.1<br>(12.1-147.9)          |
| s3            | 4   | 4/0                                                | 2/2/0                                    | 66.7<br>(35.4-88.0)           |
| s14           | 4   | 3/1                                                | 1/2/1                                    | 98.8<br>(17.5-141.9)          |
| s99           | 4   | 3/1                                                | 2/2/0                                    | 72.5<br>(43.3-152.4)          |
| s202          | 5   | 5/0                                                | 2/2/1                                    | 34.6<br>(14.4-72.6)           |

|    | A                                             | B       | C        | D       | E       | F       | G        | H       | I       | J        | K       | L       | M       | N       | O       | P        | Q       | R       | S       | T       | U        |
|----|-----------------------------------------------|---------|----------|---------|---------|---------|----------|---------|---------|----------|---------|---------|---------|---------|---------|----------|---------|---------|---------|---------|----------|
| 1  | Supplementary Table 2:                        |         |          |         |         |         |          |         |         |          |         |         |         |         |         |          |         |         |         |         |          |
| 2  | Statistical output of questionnaire analyses. |         |          |         |         |         |          |         |         |          |         |         |         |         |         |          |         |         |         |         |          |
| 3  |                                               |         |          |         |         |         |          |         |         |          |         |         |         |         |         |          |         |         |         |         |          |
| 4  |                                               |         |          |         |         |         |          |         |         |          |         |         |         |         |         |          |         |         |         |         |          |
| 5  | Group A:                                      | mono    | monoM    | oligoM  | oligoM  | oligoM  | oligoM   | oligoU  | oligoD  | oligoD   | oligoD  | oligoD  | oligoD  | oligoD  | oligoD  | oligoD   | oligoD  | oligoD  | oligoD  | oligoD  | oligoD   |
| 6  | Group B:                                      | oligo   | oligoU   | oligoU  | oligoU  | oligoU  | oligoU   | oligoU  | oligoU  | oligoU   | oligoU  | oligoU  | oligoU  | oligoU  | oligoU  | oligoU   | oligoU  | oligoU  | oligoU  | oligoU  | oligoU   |
| 7  | TFT                                           | 0.31046 | 2.00E-16 | 0.00149 | 0.00024 | 0.801   | 1.17E-05 | 0.83969 | 0.198   | 0.00627  | 0.0465  | 0.28547 | 0.686   | 0.0298  | 0.4104  | 6.51E-08 | 0.5268  | 0.3589  | 0.8915  | 0.4951  | 0.07993  |
| 8  | OS                                            | 0.4111  | 3.08E-06 | 0.5935  | 0.994   | 0.9405  | 0.0288   | 0.8831  | 0.0005  | 0.393    | 0.691   | 0.993   | 0.6816  | 0.3525  | 0.4756  | 0.0227   | 0.8535  | 0.09289 | 0.5903  | 0.9054  | 0.1603   |
| 9  | Age at dg                                     | 0.03577 | 0.00888  | 0.71112 | 0.9978  | 0.3419  | 0.1639   | 0.6226  | 0.0051  | 0.0742   | 0.4027  | 0.2074  | 0.3718  | 0.0408  | 0.1048  | 0.0135   | 0.684   | 0.6358  | 0.0404  | 0.787   | 0.0311   |
| 10 | Male sex                                      | 0.7748  | 0.0477   | 1       | 0.804   | 0.807   | 1        | 0.596   | 1       | 0.639    | 1       | 1       | 0.08133 | 0.22085 | 0.12773 | 0.46147  | 0.01704 | 0.61958 | 1       | 0.01587 | 0.35504  |
| 11 | Low risk Rai stage                            | 0.3051  | 2.50E-05 | 1       | 0.0501  | 0.792   | 1        | 0.195   | 0.384   | 0.547    | 0.684   | 0.449   | 0.449   | 0.5089  | 0.16180 | 1        | 0.4173  | 0.26034 | 0.32479 | 0.11037 | 1        |
| 12 | Intermediate risk Rai stage                   | 0.4978  | 0.0278   | 1       | 0.285   | 0.785   | 1        | 0.174   | 0.546   | 1        | 0.68    | 0.452   | 0.5897  | 0.18903 | 0.09797 | 0.48507  | 0.3998  | 0.11607 | 0.30063 | 0.36177 | 1        |
| 13 | High risk Rai stage                           | 0.7837  | 0.00239  | 1       | 0.705   | 0.589   | 0.483    | 1       | 0.512   | 0.698    | 1       | 1       | 0.5555  | 0.26009 | 0.63993 | 0.07404  | 1       | 0.399   | 1       | 1       | 1        |
| 14 | del(17p)                                      | 0.7593  | 0.00524  | 1       | 0.376   | 1       | 1        | 0.376   | 0.141   | 0.698    | 0.489   | 0.489   | 0.39973 | 0.61977 | 0.38533 | 1        | 0.2297  | 1       | 1       | 1       | 1        |
| 15 | del(11q)                                      | 0.8472  | 8.90E-13 | 0.101   | 0.00336 | 1       | 0.0102   | 1       | 0.00365 | 1        | 0.0493  | 1       | 0.0871  | 1       | 0.41498 | 0.05041  | 0.6928  | 0.06667 | 1       | 0.04501 | 0.7836   |
| 16 | Trisomy 12                                    | 0.82535 | 0.105    | 0.333   | 0.482   | 1       | 0.0802   | 0.288   | 1       | 0.379    | 1       | 0.7     | 0.1476  | 0.08543 | 0.23119 | n.a.     | 1       | 0.04958 | 0.48101 | 1       | 1        |
| 17 | Normal FISH                                   | 0.8644  | 0.466    | 1       | 0.542   | 0.571   | 1        | 0.768   | 0.789   | 1        | 1       | 0.7     | 0.40186 | 0.38276 | 0.57924 | 0.4756   | 0.57676 | 0.5748  | 0.22638 | 0.57694 | 1        |
| 18 | TP53 mutation at dg (cutoff 10% AF)           | 1       | 4.91E-13 | 0.0732  | 0.00229 | 0.796   | 0.0183   | 1       | 0.0529  | 0.585    | 0.723   | 0.0946  | 0.26873 | 0.46215 | 0.02416 | 0.62823  | 0.51481 | 1       | 1       | 1       | 1        |
| 19 | TP53 mutation at dg (cutoff 0.2% AF)          | 1       | 0.02     | 0.645   | 0.739   | 0.661   | 0.188    | 0.731   | 1       | 0.467    | 0.59    | 1       | 1       | 0.44845 | 0.73547 | 0.14153  | 1       | 0.31788 | 0.21062 | 0.1211  | 1        |
| 20 | TP53 mutation overall (cutoff 10% AF)         | 1       | 0.00296  | 0.593   | 0.476   | 1       | 0.0976   | 0.712   | 1       | 0.475    | 0.326   | 1       | 0.17489 | 1       | 1       | 0.32146  | 0.39439 | 1       | 0.15059 | 0.14554 | 1        |
| 21 | TP53 mutation overall (cutoff 0.2% AF)        | 0.54717 | 2.91E-05 | 0.378   | 0.23    | 1       | 0.091    | 1       | 1       | 0.108    | 0.317   | 1       | 1       | 1       | 0.75456 | 0.20218  | 0.68284 | 1       | 0.36364 | 0.60401 | 0.76037  |
| 22 | Living alone                                  | 0.2341  | 0.388    | 1       | 0.195   | 0.747   | 1        | 0.748   | 1       | 0.474    | 1       | 0.699   | 0.54135 | 1       | 0.436   | 1        | 1       | 1       | 0.47344 | 1       | 0.00522  |
| 23 | Children born                                 | 0.4996  | 0.4033   | 0.36643 | 0.81272 | 0.58754 | 0.10542  | 0.17368 | 0.80384 | 0.56158  | 0.14214 | 0.55567 | 0.1346  | 0.4431  | 0.3715  | 0.5539   | 0.09811 | 0.214   | 0.4801  | 0.1101  | 0.628    |
| 24 | Children born - women only                    | 0.682   | 0.4546   | 0.1585  | 0.7533  | 0.9644  | 0.1585   | 0.2389  | 0.9863  | 0.7489   | 0.2427  | 0.9819  | 0.8348  | 0.7469  | 0.671   | 0.8385   | 1       | 0.8315  | 0.04503 | 0.09431 | 0.9096   |
| 25 | Non-smokers                                   | 0.07746 | 1        | 0.0209  | 0.418   | 0.817   | 0.0375   | 0.0112  | 0.488   | 0.408    | 0.365   | 0.575   | 0.47337 | 1       | 1       | 1        | 0.52508 | 0.13788 | 0.64617 | 1       | 0.073811 |
| 26 | Smokers                                       | 0.25077 | 1        | 0.231   | 0.89    | 1       | 0.49     | 0.238   | 1       | 1        | 0.438   | 1       | 0.71222 | 0.24073 | 0.23393 | 0.22896  | 0.23095 | 1       | 1       | 1       | 1        |
| 27 | Stop-smokers                                  | 0.34116 | 0.81     | 0.168   | 0.496   | 0.618   | 0.139    | 0.108   | 0.704   | 0.563    | 0.631   | 0.783   | 0.61003 | 0.29052 | 0.42224 | 0.26462  | 1       | 0.31472 | 0.31291 | 0.58991 | 0.32812  |
| 28 | Alcohol abstinence                            | 0.1817  | 0.168    | 0.801   | 0.78    | 0.157   | 0.523    | 1       | 1       | 0.584    | 0.733   | 0.414   | 0.01379 | 0.44115 | 0.44316 | 0.30104  | 0.02479 | 0.55665 | 1       | 0.14226 | 1        |
| 29 | Alcohol occasionally                          | 0.1093  | 0.365    | 0.328   | 0.464   | 0.245   | 1        | 0.623   | 1       | 0.426    | 1       | 0.28    | 0.02748 | 0.31575 | 0.44982 | 0.20014  | 0.03277 | 0.57209 | 1       | 0.31068 | 1        |
| 30 | Alcohol regularly                             | 0.46442 | 0.609    | 0.109   | 0.246   | 1       | 0.206    | 0.237   | 0.417   | 0.537    | 1       | 0.364   | 1       | 0.48666 | 0.58875 | 0.36465  | 1       | 0.33091 | 1       | 0.04951 | 0.06651  |
| 31 | High blood pressure                           | 0.01441 | 1        | 0.134   | 0.025   | 0.0249  | 1        | 0.206   | 0.237   | 0.417    | 0.537   | 1       | 0.364   | 1       | 0.48666 | 0.58875  | 0.36465 | 1       | 0.33091 | 1       | 0.04951  |
| 32 | Low blood pressure                            | 0.61613 | 0.349    | 0.323   | 1       | 1       | 0.39     | 0.131   | 1       | 0.669    | 0.634   | 1       | 0.702   | 0.15243 | 0.34701 | 0.34149  | 0.46102 | 0.10314 | 0.33091 | 1       | 0.04951  |
| 33 | Pneumonia                                     | 0.41649 | 0.548    | 0.214   | 0.259   | 0.169   | 0.776    | 0.563   | 0.608   | 0.793    | 0.458   | 0.667   | 1       | 0.17761 | 0.25847 | 0.16755  | 0.40479 | 1       | 0.30589 | 0.57509 | 0.60365  |
| 34 | Chronic inflammation of respiratory cavities  | 0.80553 | 1        | 1       | 0.664   | 0.668   | 1        | 1       | 0.701   | 1        | 1       | 1       | 0.21178 | 0.20708 | 0.22653 | 0.20724  | 1       | 1       | 1       | 1       | 1        |
| 35 | Tuberculosis                                  | 0.17418 | 1        | 0.471   | 1       | 1       | 0.23     | 0.201   | 0.287   | 0.253    | 1       | 1       | 0.18544 | 1       | 1       | 1        | 1       | 1       | 1       | 1       | 1        |
| 36 | Chronic obstructive pulmonary disease         | 1       | 0.33     | 0.471   | 0.612   | 1       | 0.517    | 1       | 0.251   | 0.64     | 1       | 0.49    | 1       | 0.61487 | 0.61323 | 1        | 1       | 1       | 1       | 1       | 1        |
| 37 | Asthma                                        | 0.75768 | 0.85     | 1       | 1       | 1       | 1        | 0.6     | 0.612   | 0.611    | 0.432   | 0.462   | 1       | 0.46164 | 0.45565 | 0.45316  | 0.18367 | 0.30196 | 0.21575 | 1       | 0.24221  |
| 38 | Respiratory system - all                      | 0.66189 | 0.856    | 0.717   | 0.459   | 0.454   | 1        | 1       | 1       | 1        | 1       | 0.734   | 0.221   | 0.46164 | 0.45565 | 0.45316  | 0.18367 | 0.30196 | 0.21575 | 1       | 0.24221  |
| 39 | Diabetes                                      | 0.85608 | 0.91     | 1       | 0.758   | 0.753   | 0.512    | 0.522   | 0.775   | 0.775    | 0.437   | 0.682   | 0.21815 | 0.35177 | 0.35601 | 0.3429   | 0.08968 | 1       | 0.55938 | 1       | 1        |
| 40 | Hypercholesterolemia                          | 0.00469 | 0.164    | 0.738   | 0.3     | 0.619   | 1        | 0.593   | 0.00084 | 6.98E-05 | 0.0233  | 0.0549  | 0.79987 | 1       | 0.79704 | 0.80276  | 0.72811 | 0.31172 | 0.6332  | 0.59285 | 0.03671  |

