Normal complete blood count reference intervals in the Turkish population: a prospective study

Abstract
This study aims to analyze and report the normal complete blood count (CBC) reference intervals in the Turkish population. 1375 healthy people living in Manisa-Turkey, non-pregnant, with no known disease, with no pathological entity in physical examination and anamnesis, with no medicine usage in the last 15 days, stating that they and also their first degree relatives do not have any blood diseases, were included in the study. Vitamin B12, folic acid, ferritin levels were analyzed for each of the participants. CBC values of the remaining 714 people, of which 363 were women and 351 were men, were studied. Platelet (PLT), Plateletcrit (PCT), Red cell Distribution Width (RDW) levels and the neutrophil ratio were found significantly higher in females. On the other hand hemoglobin (Hb), Total Red Blood Cell count (RBC), Hematocrit (Hct), mean cellular volume (MCV), mean cellular hemoglobin (MCH), mean corpuscular hemoglobin concentration (MCHC) levels together with monocyte count and ratio were found significantly higher in males (p<0.05). World Health Organization (WHO) defines anemia as the level of Hb being less than 13g/dL for men. In our study we found this level to be 15.39±1.067(14)g/dL. WHO defines anemia as the level of Hb less than 12g/dL for women. In our study we found this level to be 13.26±1.068(12)g/dL. We found that anemia borderline of the women in our region was consistent with anemia borderline of WHO.

Keywords: anemia, turkey, hemoglobins, plateletcrit, platelet, vitamin

Abbreviations CBC, complete blood count; RDW, red cell distribution width; RBC, red blood cell count; Hct, Hematocrit; MCV, mean cellular volume; MCH, mean cellular hemoglobin; MCHC, mean corpuscular hemoglobin concentration

Introduction
There are many elements affecting the normal CBC levels, and those levels change according to the race, sex, population and the district. Studies have been performed in order to establish the reference CBC levels in many regions. As the reference levels of those studies are different from each other, we have thought to have different results also in our country. The city of Manisa (number of population in 2015: 1.380.366) is developed in terms of agriculture and industry, and allows high numbers of immigrants from other regions of Turkey (the number of immigrants between 2007-2014: 236.535 according to the data of Turkish statistical institute). We thought that a study made in the city of Manisa can form an opinion for the CBC parameters of our country.

Material and method
Study design
We have included 1375 people totally in the study, between the dates November 2011 and January 2013. 562 of them were male and 813 were female. The participants were selected from healthy people with no known disease between the ages 18-70, who were not on continuous medical treatment, did not take any medications since the last 15 days, and were not known to have hereditary blood disease in their families, with normal physical examination. All of the participants were informed, and they signed the volunteer forms.

2mL of venous blood into Ethylene daimine tetraacetic acid (EDTA)-tube and 5mL venous blood into serum tube were collected from all of the participants of the study, between 08:30 and 14:00. Those blood samples were examined within the first two hours after sample collection in the hospital laboratory. CBC and vitamin B12, folic acid, ferritin levels were analyzed for each of the participants. Since the deficiencies of vitamin B12, folic acid and/or ferritin are the most frequent reason of anemia in the population, we excluded the subjects with decreased levels of those from the study.

CBC parameters from the blood taken into EDTA-tubes were analyzed with Mindry BC-6800 device. Ferritin, vitamin B12 and folic acid levels were analyzed with DXI-800 Beckman Coulter brand device by using Beckman Coulter Syncron Systems kit.

Statistical analysis
Statistical analyses were performed by SPSS (Statistical Package for Social Sciences) for Windows 15.0.

Results
1375 people in total, from which 562 were male and 813 were female, participated in the study. Mean age of all participants was 29.61±8.6. Mean age in women was 28.65±8.52 (median: 27) and mean age in men was 30.99±8.53 (median: 27). When we excluded people with deficiency of either vitamin B12 or folic acid or ferritin, 714 people was the total number of the participants, from which 363 were female and 351 were male (Table 1).

We have examined the individuals separately according to the age, sex, place of birth and menopausal status.

While PLT, PCT, RDW, neutrophils (%) were higher in women with a statistically significant level; Hb, RBC, hematocrit, MCV, MCH, MCHC, monocyte percentage and number and eosinophile percentage and number were higher in men with a statistically significant level. No statistically significant difference was observed...
between men and women in other CBC parameters (Table 2) (Table 3).

No statistically significant difference was observed in CBC parameters between women who were born in Manisa and not born in Manisa. WBC, monocyte and neutrophil counts of men who were born in Manisa were higher with a statistically significant difference than those who were not (Table 2) (Table 3).

| Table 1 Averages of Age According to Gender |
|---------------------------------------------|
|                  | All Groups | Normal B12-Feritin-Folic Acid Group |
|                 | Women Age  | Men Age   | Total | Women Age  | Men Age   | Total |
| N               | 813        | 562       | 1375  | 363        | 351       | 714   |
| Mean            | 28.65      | 30.99     | 29.61 | 28.69      | 30.94     | 29.8  |
| Median          | 27         | 30        | 28    | 26         | 30        | 28    |
| Std. Deviation  | 8,526      | 8,537     | 8,605 | 9,241      | 8,705     | 9,045 |

| Table 2 Averages of Hemogram Value of 351 Men |
|-----------------------------------------------|
|                  | RBC | Hb  | Hct | PLT | PCT | MCV | MCH | MCHC | RDW | MPV | PDW |
| N                | 351 | 351 | 351 | 351 | 351 | 351 | 351 | 351  | 351 | 351 | 351 |
| Mean             | 5.02| 15.39| 44.69| 232 | 0.223| 89.1| 30.7| 34.4 | 13,15| 9.65| 16.3 |
| Median           | 5   | 15.4 | 44.7 | 233 | 0.219| 89.2| 30.8| 34.3 | 13   | 9.6 | 16.3 |
| Std. Deviation   | 0.408| 1.067| 3.2 | 47.86| 0.0442| 4.88| 1.95| 1.063| 0.907| 1.085| 0.563 |

| Table 3 Averages of Hemogram Value of 363 Women |
|-----------------------------------------------|
|                  | RBC | Hb  | Hct | PLT | PCT | MCV | MCH | MCHC | RDW | MPV | PDW |
| N                | 363 | 363 | 363 | 363 | 363 | 363 | 363 | 363  | 363 | 363 | 363 |
| Mean             | 4.41| 13.26| 38.82| 254 | 0.244| 88.2| 30.1| 34.1 | 13.41| 9.74| 16.3 |
| Median           | 4.39| 13.2 | 38.8 | 247 | 0.239| 88.8| 30.2| 34.2 | 13.1 | 9.7 | 16.2 |
| Std. Deviation   | 0.369| 1.068| 3.08| 56.12| 0.0488| 4.92| 1.972| 0.921| 1.778| 1.235| 0.552 |

| Table 4 Averages of Hemogram Value of 351 Men |
|-----------------------------------------------|
|                  | WBC | Neutrophil (%) | Lymphocyte (%) | Monocyte (%) | Eosinophil (%) | Basophil (%) | Neutrophil Count | Lymphocyte Count | Monocyte Count | Eosinophil Count | Basophil Count |
| N                | 351 | 351 | 351 | 351 | 351 | 351 | 351 | 351  | 351 | 351 | 351 |
| Mean             | 7.794| 58.6 | 30.12| 7.86 | 2.5  | 0.8 | 4.603| 2.303 | 0.632 | 0.19 | 0.056 |
| Median           | 7.6  | 59   | 30.4 | 6.9  | 2.1  | 0.4 | 4.4 | 2.2 | 0.5 | 0.17 | 0 |
| Std. Deviation   | 1.723| 10.15 | 7.15 | 4.96 | 1.735 | 2.49 | 1.476| 0.647 | 0.568 | 0.146 | 0.2 |

**Discussion**

There are many elements affecting the CBC reference intervals, and those levels change according to the race, sex, population and the district. That’s why the reference intervals of the hematological parameters differ from population to population.¹⁻³⁴ There have not been enough prospective studies for the reference intervals and the evaluation of the CBC parameters in our country.

The importance of our study comes from its potential to provide reference levels of hematological parameters at Turkey. We have analyzed many hematological parameters prospectively, by working on many men and women. We have also tried to obtain information

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about Turkey’s data by evaluating the individuals separately as Manisa-born and non Manisa-born.

Table 4, together with the CBC results of our study, presents the CBC results of the studies conveyed in Turkey. The most important feature of our study is its prospectively.

| Hemogram values | Terzioğlu6 | Başak7 | Kaya8 | Dilek9 | Yılmaz10 | Our study results |
|-----------------|-----------|--------|-------|--------|----------|------------------|
| RBC (10^12/L)   | 4.5±0.6  | 4.5±0.4 | 4.6±0.4 | 4.8±0.5 | 4.66±0.02 | 5.27±0.02 | 4.4±0.36 | 5.02±0.4 |
| Hb (gr/dL)      | 12.4±2.3 | 13.6±1.1 | 14.6±1.6 | 13.4±1.7 | 13.53±0.09 | 15.78±0.06 | 13.26±1 | 15.39±1 |
| Hct (%)         | 39.6±5   | 40.5±3.9 | 42.9±4.6 | 41.4±4.9 | 39.19±0.4 | 45.28±0.3 | 38.82±0.8 | 44.69±3.2 |
| PLT (10^9/L)    | 235±52  | 218±46 | 243±55 | 221±75 | 281.01±5.79 | 248.60±2.8 | 254±56.12 | 232.7±27.46 |
| PCT (%)         | 0.244±0.04 | 0.223±0.04 |
| MCV (fl)        | 88.7±4.8 | 87±4.5 | 86.7±4.6 | 87±5-84±4 | 84.01±0.47 | 85.07±0.25 | 88.2±4.92 | 89.1±4.88 |
| MCH(pg)         | 28.86±0.24 | 30.10±0.18 | 34.5±0.11 | 35.14±0.05 | 34.0±0.92 | 34.4 |
| MCHC(gr/dL)     | 13.1±1.77 | 13.15±0.9 |
| RDW             | 16.3±0.55 | 16.3±0.56 |
| MPV             | 9.08±0.32 | 8.68±0.06 | 9.7±1.23 | 9.6±1.08 |

Table 5 Comparison of our Study Results with the Other Studies Conducted in Our Country

| Hemogram values | Terzioğlu6 | Başak7 | Kaya8 | Dilek9 | Yılmaz10 | Results of our study |
|-----------------|-----------|--------|-------|--------|----------|---------------------|
| Leukocyte x10^9/L | 6.8±1.4  | 7.4±2  | 7.0±1.9 | 6.78±0.11 | 7.44±0.09 | 7.765±1.889 | 7.794±1.723 |
| Neutrophyl(%)    | 60.37±0.64 | 58.81±0.46 | 30.43±0.54 | 31.74±0.39 | 70±3.7 | 7.86±4.96 |
| Lymphocyte(%)    | 30.43±0.54 | 31.74±0.39 | 70±3.7 | 7.86±4.96 |
| Monocyte(%)      | 0.538±0.323 | 0.632±0.568 |
| Eosinophil(%)    | 2.59±0.12 | 3.04±0.09 | 1.99±1.91 | 2.5±1.73 |
| Basophil(%)      | 0.7±1.33 | 0.8±2.49 |
| Neutrophyl Count x10^9/L | 4.727±1.555 | 4.603±1.476 |
| Lymphocyte Count x10^9/L | 2.295±0.671 | 2.303±0.647 |
| Monocyte Count x10^9/L | 0.538±0.323 | 0.632±0.568 |
| Eosinophil Count x10^9/L | 0.154±0.152 | 0.19±0.146 |
| Basophil Count x10^9/L | 0.043±0.104 | 0.056±0.20 |

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Average RBC values of our study, obtained from 714 individuals between the ages 18-70 are compatible with the studies of Yılmaz et al.,10 and Tamer et al.11 But the average value obtained from our study is lower than theirs. Castro et al.12 stated that RBC values at black individuals were lower than white ones, and this may be the result of the high incidence of thalassemia and iron deficiency anemia at black population. Sirdah et al.,13 made a study between the years 2000-2008 and 94.811 people were included in the study. But enough information was not given about the selection of healthy individuals and the ones with low levels of ferritin were included in the study.10-13

Roshan et al.,14 gave Hb values as 11.83±1.01 in women and 14.27±1.13 in men. When compared with the result of our study, their average Hb values and reference intervals were lower both in men and women. We assume that the difference in Hb values between the results of Roshan et al.,14 and us is due to the differences in geographical location, health, nutrition, gender and smoking status. Besides Roshans et al.,14 did not exclude the individuals with vitamin B12, folic acid and iron deficiencies from their study, which are actually the most common causes of anemia in the world.14

Sirdah et al.,13 found Hb values as 15.09±1.09 in smoking men, 14.74±1.09 in non-smoking men and 12.36±1.18gr/dL in non-smoking women. Sirdah et al.,13 found that Hb values of non-smokers are higher than the Hb values of smokers with a statistically significant difference.15 While WHO accepts the value under 13gr/dL as anemia in men, we have obtained this value to be 15,39±1,067gr/dL in our study (14gr/dl). WHO accepts Hb value under 12gr/dL as anemia in women, we have obtained this value to be less than 13,26±1,068(12) gr/dL, which is compatible with the result of WHO.

In our study, the average of the Hct values obtained from all of the participants aging between 18-70 were compatible with the studies of Tuncer et al.,16 Tikly et al.,16 Kelly & Munan.17

Platelet values in women were found to be significantly higher than the values in men (p = 0.012) in our study. Although our normal values were compatible with the values of Tamer et al.,11 and Tuncer et al.,15 our average value was higher than Tamer et al.,11 average value.12,13 Researchers stated that platelet numbers were higher in women than men. This difference becomes clearer in the premenopausal period. Our study was also compatible with these results. In addition, we have found the number of platelet to be higher in women in menopause than women not in menopause.

When it comes to the comparison of average WBC values, Tikly et al.,16 stated that the number of WBC count was lower in black individuals than white ones, and they also stated that the reason of that difference was not obvious. Kuevakoje et al.,15 did not find statistically significant difference in WBC values between men and women. Sirdah et al.,13 found that while WBC values of nonsmoking women were significantly higher than the values of nonsmoking men, it was also significantly higher in smoking men than in nonsmoking men (p<0.001). We didn’t evaluate the individuals separately as smoker or nonsmoker in our study.16,18,13

Although there are some studies stating that WBC numbers are decreasing with the rise in altitude, there are also other studies that don’t support this view. Rana et al.,19 found WBC numbers higher in the adults of black race. The reason of this is attributed to non-exclusion of the pregnant women from the participants.19 We have not included the pregnant women in our study, so the limits of our normal value seems to be higher than the results of those studies. We have also compared the WBC values between men and women and we saw that WBC values were higher in women. But we did not observe a statistically significant difference in WBC values between the women in menopause and not in menopause.

The averages of leukocyte, PLT, RBC- Hemoglobin-Hematocrit-MCV-MCHC values in different ethnicities and %90 confidence interval of these values were shown together with our results in Table 6 and 7.12,18,19 It is seen that WBC, % neutrophil, % lymphocyte, % monocytes, % eosinophile and platelet numbers are found higher in our study than the studies conducted on other ethnicities.

Consequently, with this prospective study, we have obtained data about normal CBC reference intervals in our region and in Turkish population by evaluation of 714 healthy participants with normal vitamin B12, folic acid and ferritin levels.

The most statistically significant result of this study is that, while WHO accepts values under 13gr/dL as anemia for men, we have found this value to be 15.39±1.067(14)gr/dL in our study. WHO accepts values under 12gr/dL as anemia in women, and we have found this value to be 13.26±1.068(12)gr/dL in our study. So our results are compatible with WHO’s data. Performing studies with larger number of healthy volunteers who live in different regions will be more useful.

Table 6 Leucocyte-platelet count(x10^9/L) and percentile intervals(%5-95) in different ethnicities

| CBC Reference Intervals | Men                          | Women                          |
|-------------------------|------------------------------|--------------------------------|
|                         | Caucasian | African | European | African | African | Results of our study |
| Number                  | n = 100   | n = 65  | n=351    | n = 100 | n = 50  | n=363              |
| Leukocyte (x10^9/L)     | 5.7       | 4.5     | 7.99     | 5.7     | 5       | 7.7                |
| %90 CI                  | (3.6–9.2) | (2.8–9.5)| (5.4–10.6) | (3.5–10.8) | (3.2–7.8) | (5.1–11.1) |
| Neutrophyl(x10^9/L)     | 3.2       | 1.95    | 4.6      | 3.6     | 2.4     | 4.7                |
| %90 CI                  | (1.7–6.1) | (0.9–4.2)| (2.5–7.4) | (1.7–7.5) | (1.3–4.2) | (2.7–7.9) |
| Lymphocyte(x10^9/L)     | 1.7       | 1.8     | 2.3      | 1.8     | 2       | 2.2                |
| %90 CI                  | (1.0–2.9) | (1.0–3.2)| (1.0–3.5) | (1.2–3.4) | (1.1–3.6) | (1.4–3.4) |
| Monocyte(x10^9/L)       | 0.34      | 0.29    | 0.6      | 0.3     | 0.31    | 0.53               |
| %90 CI                  | (0.18–0.62)| (0.15–0.58)| (0.14–0.61)| (0.16–0.59)| (0.15–0.39)| (0.3–0.8) |
| Eosinophil(x10^9/L)     | 0.12      | 0.19    | 0.12     | 0.13    | 0.1     | 0.15               |
| %90 CI                  | (0.03–0.48)| (0.02–0.79)| (0.04–0.44) | (0.03–0.33)| (0.02–0.41)| (0.0–0.4) |
| PLT(x10^9/L)            | 218       | 232     | 246      | 236     | 207     | 254                |
| %90 CI                  | (143–332) | (115–290)| (169–358) | (149–374) | (125–342) | (171–346) |

CI: Confidence interval

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### Table 7 RBC-Hemoglobin-Hematocrit-MCV-MCHC and percentile intervals(5%-95) in different ethnicities

| CBC | Reference Intervals | American Results of our study | African Results of our study | South Africa Results of our study | Hispanic Results of our study | Asian Results of our study | American Results of our study | African Results of our study | South Africa Results of our study | Hispanic Results of our study | Asian Results of our study |
|-----|---------------------|-------------------------------|-----------------------------|-----------------------------------|-------------------------------|-----------------------------|-------------------------------|-------------------------------|---------------------------------|-------------------------------|-----------------------------|
| RBC (S) Min %90 CI (x10^7/L) | 4.5–5.9 | 5.02 ±0.4 | 4.5–6.1 | 3.2–5.8 | 4.14–5.68 | 4.06–5.97 | 4.0–5.2 | 4.41 ±0.36 | 3.42–5.44 | 3.0–5.3 | 3.71–5.06 | 3.66–5.05 |
| Hemoglobin %90 CI % | 13.5–17.5 | 15.39 ±1 | 12.3–17.3 | 10.3–16.7 | 13.5–17.0 | 12.2–16.9 | (12.0–16.0) | 13.26 ±1 | 9.1–14.9 | 9.0–15.2 | 10.2–14.8 | 14.9 |
| Hematocrit %90 CI | 41–53 | 44.69±3.2 | 39–52 | 31–52.5 | 38.8–49.5 | 36.7–49.4 | 36–46 | 38.82±3.08 | 28–44 | 27.3–47.2 | 31.0–44.1 | 32.2–43.8 |
| MCV Min %90 CI | 80–100 | 89,1±4.88 | NA | NA | 82.3–98.4 | 69.9–99.8 | 80–100 | 88.2 ± 4.92 | NA | NA | 1.7–7.5 | 1.4–6.5 |
| MCHC g/dL Min %90 CI | 34.4 ±1.06 | NA | NA | NA | 32.4–35.8 | 31.8–36.0 | NA | 34.1±0.92 | NA | NA | 32.3–35.7 | 32.3–35.8 |

Cl: Confidence interval, NA: Not Available

### Ethical approval

It is received from the Ethics Committee of the Manisa Celal Bayar University in Manisa/Turkey. All subjects signed an informed consent and the study was conducted in accordance to the principles of the Declaration of Helsinki.

### Declaration of authorship

EAK, ÜE designed the study. EAK, ÜE and CO performed data acquisition. EAK, CO analyzed and interpreted the data. EAK and UE drafted the manuscript. All co-authors critically reviewed the manuscript and gave their final approval of the version of the manuscript to be published.

### Competing interests

The author declares that there are no conflicts of interest.

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