PRACTICAL ADVICES FOR PHYSICIANS

Understanding sensitivity, specificity and predictive values

Razumevanje osetljivosti, specifičnosti i prognostičkih vrednosti

Miodrag Stojanović*, Marija Apostolović*, Dijana Stojanović*, Zoran Milošević*, Aleksandra Toplaović*, Vesna Mitić Lakušić†, Mlađan Golušović‡

*Faculty of Medicine, University of Niš, Niš, Serbia; †Clinical Center Podgorica, Podgorica, Montenegro; ‡Clinical center Niš, Niš, Serbia

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Introduction

Modern medicine and medical science have experienced a change in knowledge and dramatic increase in technology. Nowadays “Hi-Tech” tests are widespread and doctors can hear a statement: “This new test is very reliable, it is the most sensitive test at the market, you should use it”, practically every day. The basic idea for using diagnostic tests aims to easier making the diagnosis of a disease and so enable appropriate treatment. How to know if a test is useful? Will a test point out to a doctor whether a patient is ill or not?

Due to the fact that the doctors and medical research workers are not so common with the statistical methods, the aim of this paper was to explain the basics of sensitivity, specificity and predictive values in a very simple way, using simple logic formulas, and presenting the samples from medical practice.

Diagnostic test and the gold standard

Diagnostic tests are all the tests that physician can use in the process of making the diagnosis of a particular disease. It is a procedure performed to confirm or determine the presence of a disease in a person suspected of having the disease, usually following the report on signs and symptoms, or based on the results of other medical tests 1.

The ideal test should correctly identify all tested people with or without disease with 100% of accuracy, which is practically impossible. Traditionally, to help understanding sensitivity, specificity and predictive values, the best method for explanation is based on 2 x 2 contingency table. Suppose a population of 1,000 people, 100 of them have a disease X, 900 do not have the disease according to the gold standard results (Table 1). Here, the rows represent the screening test results and the columns the true condition of a person according to the gold standard. Screening test is used to identify 180 people with the disease (a + b).
Sensitivity

Sensitivity of a clinical test represents test ability to correctly identify people with illness (a) within all people with illness (a + c). It is a proportion of people with disease who positive, expressed in percentages. Sensitivity as a fixed test characteristic provides a true positive rate.\(^5\,\,6\).

\[
\text{Sensitivity} = \frac{\text{True positives}}{\text{True positives} + \text{False negatives}}
\]

If we apply screening test to our hypothetical population and receive that 80 of the 100 people with disease X test positive, than the sensitivity of this test is 80/100 or 80% (Table 1). A test with 80% of sensitivity detects 80% of true ill patients, while 20% (false negative) will not be detected.

Specificity

The specificity of a clinical test represents test ability to correctly identify people without illness (d) within all people free from illness (b + d). It is a proportion of people without disease who test negative. Specificity is also a fixed characteristic of the test and represents true negative rate.\(^5\,\,8\).

\[
\text{Specificity} = \frac{\text{True negatives}}{\text{True negatives} + \text{False positives}}
\]

This hypothetical population (Table 1) demonstrates that 800 of the 900 people without disease X test negative, so specificity of this screening test is 800/900 or 89%. Practically, this test detects 89% of healthy people but 11% will be assumed as false positive.

For a test to be accurate, both sensitivity and specificity should be high. When measuring sensitivity, we only calculate those people with disease. High sensitive test detects a high percentage of positive cases while missing few. Also, a negative result would suggest the absence of disease according to test with high sensitivity. On the contrary, specificity highlights negative test results. A highly specific test is good for detection of a disease if a person tests positive, likewise it does not falsely diagnose disease when none is present.

It is worthy to mention that specificity and sensitivity of a quantitative test depend on a cut-off value. This is the value which determines the limit between positive and negative test results. In a situation when the cut-off is reduced, most people with the disease will be correctly identified, but at the same time the number of false positives will be increased. Raising the cut-off value will show more false negatives, but will reduce the number of false positives.\(^5\,\,10\). Practically, sensitivity and specificity are inversely proportional, as one increases the other decreases and vice versa.

The lack of adequate education about interpreting test results (especially relevant to tests that may have minor and insignificant findings) may lead to misdiagnosing.\(^11\) It is known that in two cases if test result doesn't correspond to the real situation an error has occurred. Therefore it is essential to pay attention to false positive and false negative test results. These errors are closely associated with the terms of type I and type II errors in hypothesis testing. In hypothesis testing "null hypothesis" matches to the natural state (in our situation people who are free from disease). As opposed to the null hypothesis there is an "alternative hypothesis" which corresponds to the ill people. Type I error, also known as "α" error appears when we reject null hypothesis which is actually correct.\(^12\,\,13\). Type I error corresponds to the false positive results. Type II error or "β" error appears when accepting null hypothesis, when actually it is not really true.\(^12\,\,13\). This error corresponds to false negative test results. It depends on a situation in which a false result is more undesirable. Minimizing false positives and false negatives at the same time maximizes sensitivity and specificity. Generally, it is not benign to tell someone after testing that he has a serious disease (false positive) when he does not really have the same (HIV for example). Moreover, it is inexcusable to overlook a disease when it really exists (false negative).

Sensitivity and specificity do not depend on the disease prevalence.\(^14\) They are conditional on the patient either having or not having a disease and represent the power of a diagnostic test to discriminate between those with and without disease. When a patient has a positive test result, does it actually mean that he/she has disease or not? Sensitivity and specificity cannot answer such a question, thus it is worthy to know predictive values.

**Predictive values**

The real questions to be answered are the following: “What is the probability that a person with a positive test results will have the disease? Also if a person has a negative test, what is the likelihood that he is healthy?” These questions refer to what’s called the “predictive values”. Therefore, the mission of the clinician is to determine the likelihood of a disease present given a positive test (positive predictive value – PPV), or the likelihood of a disease absent given a negative test (negative predictive value – NPV).

**Positive predictive value**

The positive predictive value or precision rate is defined as a proportion of people with a positive test result (a) who are actually ill \((a + b)\).\(^5\,\,9\). It is calculated by the formula:

\[
\text{Positive predictive value} = \frac{a}{a + b}
\]
In our population from 180 (a + b) people who test positive, 80 (a) of them actually have a disease X. PPV in this case is 80/180 or 44%. This practically means that a person who tests positive has a 44% likelihood of having a disease X. Less false positives (higher specificity) signifies a higher PPV in the observed population.

**Negative predictive value**

The negative predictive value is defined as a proportion of people with a negative test result (d) who actually do not have disease (b + c) \(^5, 8\). The formula for this measure is:

\[
\text{Negative predictive value} = \frac{\text{True negatives}}{\text{True negatives} + \text{False negatives}}
\]

We can see from our example (Table 1) that out of 820 people disease free, 800 people test negative. That basically means, a person who tests negative has 800/820 = 0.97 or 97% likelihood of not having a disease. NPP of enormous 99% indicates that a negative screening test actually dismisses disease. On the other hand, a very low PPV (only 44%) indicates that every second ill patient is diagnosed wrongly according to this test. A highly sensitive test (small number of false negatives) will have a higher NPP in the observed population.

In clinical practice, the value of a test result for an individual patient depends on the prevalence of tested population (the proportion of the population that has the disease at a given time) \(^5, 16\). We should expect that for any given population, as the prevalence of a disease increases, the test PPV will also increase and vice versa, as disease prevalence in the population being tested decreases, the PPV of that test will also decrease, while the NPP will increase. To help you understand these relationships we will illustrate this through the following example. Imagine that we now apply the same screening test we used before (Table 1) to another population where the disease prevalence is 50%, respectively 500 sick patients out of 1,000 examined (Table 2). Since this is the same test, sensitivity and specificity remain the same 90% and 80%, respectively. According to this new scenario, PPV is 450/550, or 82%, whereas the NPP is 400/450, or 89%. In the case when we repeat the same test to some other population with different disease prevalence, we will notice that the PPV increases with the increasing disease prevalence while the NPP decreases in the same situation (Table 3). Though the sensitivity and specificity remain the same, the PPV has changed remarkably.

<table>
<thead>
<tr>
<th>Test</th>
<th>Disease present (n)</th>
<th>Disease absent (n)</th>
<th>Total (n)</th>
</tr>
</thead>
<tbody>
<tr>
<td>Positive</td>
<td>450</td>
<td>100</td>
<td>550</td>
</tr>
<tr>
<td>Negative</td>
<td>50</td>
<td>400</td>
<td>450</td>
</tr>
<tr>
<td>Total</td>
<td>500</td>
<td>500</td>
<td>1000</td>
</tr>
</tbody>
</table>

Prevalence = 500/1000 = 0.5; sensitivity = 450/500 = 0.9; specificity = 400/500 = 0.8; PPV = 450/550 = 0.82; NPV = 400/450 = 0.89; PPV – positive predictive value; NPV – negative predictive value.

<table>
<thead>
<tr>
<th>Prevalence (%)</th>
<th>Positive predictive value (%)</th>
<th>Negative predictive value (%)</th>
</tr>
</thead>
<tbody>
<tr>
<td>5</td>
<td>0.19</td>
<td>0.99</td>
</tr>
<tr>
<td>10</td>
<td>0.44</td>
<td>0.97</td>
</tr>
<tr>
<td>50</td>
<td>0.82</td>
<td>0.89</td>
</tr>
<tr>
<td>70</td>
<td>0.90</td>
<td>0.77</td>
</tr>
<tr>
<td>90</td>
<td>0.96</td>
<td>0.47</td>
</tr>
</tbody>
</table>

The utility of predictive values is limited because the statistics is determined by sensitivity and specificity of a test, as well as with the prevalence of disease which can vary \(^7\). In general, specificity has more impact on a positive predictive value in the case of low disease prevalence \(^10\). Constancy of sensitivity and specificity are an important feature of a test when using in similar patients and similar settings. Predictive values although associated with sensitivity and specificity will change with the prevalence of target disease (Figure 1).
It is vital to note while comparing two populations that predictive values are only significant if the disease prevalence is the same in both populations. They are used along with the specificity and sensitivity when the prevalence in the target population is known. In the case of unknown prevalence, the sensitivity and specificity are the primary measurement used to evaluate the accuracy of a test 19.

Conclusion

To enable interpretation of diagnostic test accuracy it is necessary to understand the concepts of sensitivity, specificity and predictive values. These calculations require a design of the two-by-two table. Sensitivity and specificity of a test have limited clinical usefulness of the disease likelihood in an individual patient. Although sensitivity and specificity are not under the influence of the disease prevalence they can be affected by the differences in disease characteristics. In everyday clinical practice knowing the predictive values is more significant for measuring diagnostic accuracy. So, next time you hear about a new reliable test, ask yourselves what gold standard is performed, what is the disease prevalence, and most important of all, what are the sensitivity, specificity, positive and negative predictive values? If the answers are satisfactory, you can recommend this test for use.

References


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