

thaggurinig abiriban,
so as we will discuss in writing a
MET protein over expression in
non-small cell lung cancer.
So, coming to the methalteration
in non-small cell,
so basically it is divided in four groups.
MET exon 14 skipping mutation,
MET amplification,
MET protein over expression
and MET fusion.
So, what are the clinical significance
of each of this?
So, MET exon 14 skipping can be seen
up front in treatment naïve patients
as well as as a resistant pathway.
After EGFR and LTK exposure in respect to patients
and we have whatever the available TKI,
like cap-mountain event depotin,
these are most effective in this subgroup of the patients.
Then next is the MET amplification.
Again MET amplification is commonly seen
as a resistant pathway after exposure to EGFR and LTK.
Rarely 1 to 5% of the cases of methalteration
present as a front MET amplification.
Again, these subgroups are generally considered
as a little resistant to the available MET TKI,
like cap-mountain event depotin,
though they can be effective,
but the magnitude benefit is significantly lower
as compared to the exon 14 skipping mutation.
Then next is the MET overexpression.
These are generally seen up front
and also can be seen as a resistant pathway
after exposure to different TKIs.
Overall the MET overexpression
till few years back,
there was no any effective therapy
and these are just considered
as a poor prognostic marker.
Last are the MET fusions which are very rare.
0.2.3% cases can be seen in non-small cell lung cancer,
but these are TKI sensitive group.
If these are positive,
definitely they can go ahead with the starting TKI as a front therapy.
So today's we will be discussing the MET overexpression
population in non-small,
non-smokers easy for wild type non-small cell lung cancer patients.
So these are results from luminosity
in Asian subgroup of the patients.
So basically,
TELISOV is antibody drug conjugate
where it has a payload in form of MMA, MMA,
MMA which is mono-methyl-ore-statin E,
which is a micro-tubule inhibited cytotoxic agent.
So approximately 25% of the patients
which are non-smokers easy for wild type NSCLC express this
MET protein by ISC.
And the significance is that these are considered as a

poor prognostic group lead to deterioration of the overall survival in globally including Asian race.

In the in the luminosity trial, which is the phase 2 trial, two stage, in the stage 1 they identified the optimal subgroup for C protein overexpressing non-small cell population to treat with TELISOV and the stage 2 which is expansion group, they studied the efficacy and safety population.

So overall, this is the study design where you can see that they included patients who are more than 18 years who had advanced metastatic NACLC and they had C MET protein overexpression. This C MET protein overexpression was defined as a high or intermediate. So any NACLC patients who are having more than 25% expression of MET protein by ISC and having a 3 plus staining will be considered as a MET overexpressure.

And out of those who are having more than 50% of ISC staining will be considered as a high and between 25 to 49% will be considered as an intermediate expression.

And these patients included those who had received at least two lines of or less than two lines of therapies.

So in the stage 1 they included different subgroup, a non-scomers EGF or wild type, non-scomers EGF are mutated and scomas.

So in this they found non-scomers EGF are mutated and scomas was not eligible for further testing only in the phase 2.

They continued with the non-scomers EGF are wild type.

The primary endpoint of the study was over a response rate and the secondary.

This is controlled at duration of response, PFS and OS.

So this is the demographic of overall population where almost 70% were male population, around 30% were Asian population and in 20% population had upfront brain mates made over expression in high and intermediate were approximately 50-50% and almost treatment details you can see that almost 97.5% of population received platinum-based therapy and around 80% population has received immunotherapy drug.

So coming to the first overall response rate.

So overall there was around 28.6% patient has shown response in this population which is around third line or second line and between these two divide,

see met high expression has shown 34.6% response and intermediate group has shown 20 to 9% response and these responses were durable in the range of 8 months and at more than 6 months those who has response had a durable response in the range of 56%.

Again those who had high expression had a higher rate of durable response 63% and coming to the side effects were very far in edema, sensory neuropathy, fatigue and in some cases ILD.

So coming to the out of these the Asian subgroups so out of these 150 patients and 160 patients there was 57 patients who were Asian and out of these only efficacy was available in 48 patients again in these 48 patients, around 40-15% cases has brain mates and treatment details again you can see

that almost major group the patient has received platinum based or immun check inhibitor therapy.
So this is the patient disposition almost treatment median duration of treatment was 4.7 months.
Those patients who discontinued drug almost 50% patient had progression plus two patients has clinical progression and two patients had three patients had withdrawn the consent and among the patients who discontinued study almost half of the population was died.
So this is again response rate in Asian population almost similar to the global population with 35% and those who are having higher expression had 46.2% response rate.
This is the PFS almost it was in third line around 5.5 month was the PFS and the overall survival was in the range of 17 to 18 months.
And those who had a higher expression had a longer overall survival would be 25 to 4%.
Again the toxicity was similar across the globe in Asian population with the most common side effects were peripheral neuropathy, pneumonia and fatigue.
So conclusion so treatment with TELISOV has shown similar outcome in Asian subgroup and the side effect profile was manageable and similar to global population and for this further the pastry study is going on where they compare TELISOV with the dosi taxel.
Thank you.